

Exhibit 3

**THE UNITED STATES DISTRICT COURT
FOR THE MIDDLE DISTRICT OF PENNSYLVANIA**

TAMMY KITZMILLER, et al.,

Plaintiffs,

v.

DOVER AREA SCHOOL DISTRICT and
DOVER AREA SCHOOL DISTRICT
BOARD OF DIRECTORS,

Defendants.

Case No. 04-CV-2688

Hon. Judge Jones

**DECLARATION OF
DR. MICHAEL BEHE**

I, Michael Behe, make this declaration pursuant to 28 U.S.C. § 1746 and based on my personal knowledge.


1. I am an adult resident of the State of Pennsylvania, a citizen of the United States, and a Professor of Biological Sciences at Lehigh University in Bethlehem, Pennsylvania. I make this declaration in support of Defendants' Motion for Summary Judgment.

2. I have been retained by Defendants to provide expert testimony in this case. Attached to this declaration as Exhibit A is a true and accurate copy of my Federal Rule of Civil Procedure 26 Disclosure of Expert Testimony. Attached to this declaration as Exhibit B is a true and accurate copy of my Rebuttal Analysis of Kenneth Miller's Statement. Attached to this declaration as Exhibit C is a true and accurate copy of my Rebuttal Analysis of Kevin Padian's Statement.

3. Exhibits A, B, and C contain a complete and accurate statement of all of my opinions expressed in this case to date and the basis and reasons for them; the data or other information I considered in forming the opinions; any exhibits to be used as a summary of or support for these opinions; my qualifications as an expert, including a list of all publications authored by me within the preceding ten years; and the compensation I will receive for my study and testimony. If called to testify at trial in this case, I will testify as to the opinions contained in these Exhibits.

I declare (or certify, verify, or state) under penalty of perjury under the laws of the United States of America that the foregoing is true and correct.

Executed on this 7 day of July, 2005.



Michael J. Behe

Federal Rule of Civil Procedure 26 Disclosure of Expert Testimony

of

Michael J. Behe

Professor of Biological Sciences

Lehigh University

Bethlehem, Pennsylvania

1) My qualifications as an expert witness:

PROFESSIONAL EXPERIENCE

9/97-present	Professor of Biological Sciences, Lehigh University
1996-present	Fellow, Discovery Institute's Center for Science and Culture
6/95-8/97	Associate Professor of Biological Sciences, Lehigh University
9/85-6/95	Associate Professor of Chemistry, Lehigh University
7/89-12/89	Visiting Associate Professor of Biochemistry, Hershey Medical Center/Penn State
9/82-8/85	Assistant Professor of Chemistry, City University of New York, Queens College
11/78-9/82	Jane Coffin Childs Fund Postdoctoral Fellow at the National Institutes of Health (Gary Felsenfeld, advisor)
9/74-10/78	National Research Service Award Predoctoral Fellow at the University of Pennsylvania (Walter Englander, mentor)

EDUCATION

Ph. D.	Biochemistry, 1978. University of Pennsylvania, Philadelphia, Pennsylvania.
B.S.	Chemistry, 1974. Drexel University, Philadelphia, Pennsylvania.

PUBLICATIONS IN THE LAST TEN YEARS

Behe MJ, Snoke DW. 2004. Simulating evolution by gene duplication of protein features that require multiple amino acid residues. *Protein Sci* 13:2651-2664.

Behe, M.J. 2004. Rationalism and reform: A review of Who Rules in Science? by James Robert Brown, *First Things*, August/September, pp. 75-79.

Behe, M.J. 2004. "Irreducible Complexity: Obstacle to Darwinian Evolution." In *Debating Design: from Darwin to DNA*, Ruse, M. and Dembski, W.A., eds., Cambridge University Press, pp. 352-370.

Behe, M.J. 2003. "Design in the Details: The Origin of Biomolecular Machines." In *Darwinism, Design & Public Education*, Campbell, J.A. and Meyer, S.C. eds., Michigan State University Press, pp. 287-302.

Behe, M.J. 2003. "Darwin's Black Box." In *Faith, Freedom, and the Future: Religion in American Political Culture.* Dunn, C.W. ed., Rowman and Littlefield, pp. 87-100.

Behe, M.J. 2003. "The Modern Intelligent Design Hypothesis: Breaking Rules." In *God and Design: The Teleological Argument and Modern Science*, Neil Manson, ed., Routledge, pp. 277-291.

Behe, M.J. 2002. The challenge of irreducible complexity. *Natural History* **111**, 74.

Behe, M.J. "Molecular Machines: Experimental Support for the Design Inference." In Robert T. Pennock (ed.) *Intelligent Design Creationism and Its Critics: Philosophical, Theological and Scientific Perspectives*. 2001. Cambridge, MA & London, England: The MIT Press. pp. 241-256.

Behe, M.J. 2001. Reply to My Critics: A Response to Reviews of Darwin's Black Box: The Biochemical Challenge to Evolution, *Biology and Philosophy* **16**, 685-709.

Behe, M.J. 2001. Abstract, Protein Society meeting. A Population Genetics Model for the Number of Generations Needed to Develop a New Protein Binding Site. *Protein Science* **10**, 65.

Behe, M.J. 2001. Review of "Extreme Functional Sensitivity to Conservative Amino Acid Changes on Enzyme Exteriors." *ChemTracts—Biochemistry and Molecular Biology* **14**, 152-158.

Behe, M.J. 2001. A review of *Finding Darwin's God*, *National Catholic Bioethics Quarterly* **1**, 277-278.

Behe, M.J. 2001. The Modern Intelligent Design Hypothesis: Breaking Rules, *Philosophia Christi* **3**, 165-179.

Behe, M.J. 2001. "Faith and the Structure of Life." In *Science and Faith*, Gerard V. Bradley and Don DeMarco, eds. St. Augustine's Press, South Bend, Indiana, Chapter 2.

Behe, M.J. 2001. "Darwin's Breakdown: Irreducible Complexity and Design at the Foundation of Life." In *Signs of Intelligence: Understanding Intelligent Design*, William A. Dembski, ed. Brazos Press, Grand Rapids, Michigan, Chapter 7.

Behe, M.J. 2001 Darwin's Black Box, Navrat Domu, Prague. (The Czech translation of Darwin's Black Box.)

Behe, M.J. 2001 Darwin's Black Box, KCC, Seoul. (The Korean translation of Darwin's Black Box.)

Behe, M.J. 2000. "Evidence of Design in Biology." In M.J. Behe et al., *Science and Evidence for Design in the Universe*, Ignatius Press, San Francisco, Chapter 3.

Behe, M.J. 2000. "Answering Scientific Criticisms of Intelligent Design." In M.J. Behe et

al., *Science and Evidence for Design in the Universe*, Ignatius Press, San Francisco, Chapter 4.

Behe, M.J. 2000. Conservatives, Darwin, and Design: An Exchange. *First Things*, no. 107, November, pp. 28-29.

Behe, M.J. 2000. Evidence of design in biology. *Providence* 5, 23-32.

Behe, M.J. 2000. Review of "Tolerance of Arc repressor to multiple-alanine substitutions." *ChemTracts—Biochemistry and Molecular Biology* 13, 409-414.

Behe, M.J. 2000. Self-Organization and Irreducibly Complex Systems: A Reply to Shanks and Joplin. *Philosophy of Science* 67, 155-162.

Behe, M.J. 2000. Evolution, Religion, and the National Academy of Sciences, *Ethics and Medics* 25, 1-3.

Behe, M.J. 1999/2000. Darwin's Hostages. *The American Spectator* 32, December-January, pp. 32-36.

Behe, M.J. 1999. Review of Nature's Destiny by Michael J. Denton, *Persp. Biol. Med.* 43, 151-153.

Behe, M.J. 1999. Review of God After Darwin: A Theology of Evolution by John Haught, *Metaviews*, December, www.meta-list.org.

Behe, M.J. 1999. Teach evolution without bias. *ExploreZone* webzine, September 7. explorezone.com/archives/99_09/07_opinion_kansas_behe.htm. (This was a companion piece to one written by Bruce Alberts, president of the National Academy of Sciences. explorezone.com/archives/99_08/30_opinion_kansas.htm)

Behe, M.J. 1999. Darwin's Black Box, Central Compilation and Translation Publishers, Beijing. (The Chinese translation of Darwin's Black Box.) (<http://cctpbook.com>)

Behe, M.J. 1999. Teach Evolution—And Ask Hard Questions. *New York Times*, August 13, p. A21.

Behe, M.J. 1999. Comment on Denis Lamoureux's Essays. In Johnson, P.E. and Lamoureux, D.O., Darwinism Defeated? Regent College Press, Vancouver, pp. 103-108.

Behe, M.J. 1999. Darwin's Breakdown. *Touchstone* 12, 39-43.

Behe, M.J. 1999. The God of Science: The Case for intelligent Design. (Review of Tower of Babel: The Evidence Against the New Creationism by Robert Pennock), *The Weekly Standard*, June 7, pp. 35-37.

- Behe, M.J. 1999. Review of The Fifth Miracle: The Search for the Origin and Meaning of Life by Paul Davies, *First Things*, June/July, pp. 42-45.
- Behe, M.J. 1998. Intelligent Design as an Alternative Explanation for the Existence of Biomolecular Machines. *Rhetoric & Public Affairs* 1, 565-570.
- Behe, M.J. 1998. Intelligent Design Theory as a Tool for Analyzing Biochemical Systems, in Mere Creation, Dembski, W., ed., InterVarsity Press, Carol Stream, Illinois, pp. 177-194.
- Behe, M.J. 1998. Tracts of separated, alternating, and mixed adenosine and cytidine residues in the genomes of prokaryotes and eukaryotes. *DNA Sequence* 8, 375-383.
- Behe, M.J. 1998. Tulips and Dandelions: A review of A Case Against Accident and Self-Organization. *Books and Culture*, September/October, pp. 34-37.
- Behe, M.J. 1998. Darwin's Black Box, Seidosha, Japan. (The Japanese translation of Darwin's Black Box.)
- Behe, M.J. 1998. Darwin's Black Box, Ankara. (The Turkish translation of Darwin's Black Box.)
- Behe, M.J. 1998. Embryos and Evolution (letter to the Editor). *Science* 281, 348.
- Behe, M.J. 1998. Dogmatic Darwinism, *Crisis*, June, pp. 32-36
- Behe, M.J. 1998. Randomness or Design in Evolution, *Ethics and Medics* 23, 3-4.
- Behe, M. J. 1998. Darwinism and Design. In Brungs, R. and Postiglione, M., eds. Creation and Evolution, ITEST Faith/Science Press, St. Louis, Missouri, pp. 1-25.
- Behe, M.J. 1998. "Darwin Controversy: A Commentary on John O. McGinnis's essay", *National Review*, March 9, pp. 53-54.
- Behe, M.J. 1998. Darwin's Black Box: The Biochemical Challenge to Evolution, Touchstone- Simon & Schuster, New York (the paperback edition).
- Behe, M.J. 1998. A review of Adrian Desmond's Huxley: From Devil's Disciple to Evolution's High Priest, *National Review*, February 9, 54-56.
- Mahloogi, H. & Behe, M.J. 1997. Oligoadenosine tracts favor nucleosome formation. *Biochem. Biophys. Res. Comm.* 235, 663-668.
- Behe, M.J. 1997. De Zwarte Doos van Darwin, Ten Have, Netherlands. (The Dutch translation of Darwin's Black Box.)

Behe, M.J. 1997. A Caixa Preta de Darwin, Jorge Zaha Editor, Brazil. (The Portuguese translation of Darwin's Black Box.)

Behe, M.J. 1997. Dragging Darwin In: A review of Matt Ridley's The Origins of Virtue: Human Instincts and the Evolution of Cooperation, *National Review*, June 2, pp. 52-54.

Behe, M.J. 1997. The Sterility of Darwinism, *Boston Review*, Feb-March, p.24.

Behe, M.J. 1996. Darwin Under the Microscope, *New York Times*, October 29, p. A25.

Behe, M.J. 1996. Clueless at Oxford: A review of Richard Dawkins' Climbing Mount Improbable, *National Review*, Oct. 14, 1996, pp. 83-85.

Behe, M.J. 1996. Darwin's Black Box: The Biochemical Challenge to Evolution, The Free Press, New York.

Agarwal, S. & Behe, M.J. 1996. Nonconservative mutations are well-tolerated in the globular region of yeast histone H4. *J. Mol. Biol.* **255**, 401-411.

Behe, M.J. 1995. An overabundance of long oligopurine tracts occurs in the genome of simple and complex eukaryotes. *Nucleic Acids Res.* **23**, 689-695.

Puhl, H.L. & Behe, M.J. 1995. Poly[dA]-poly[dT] forms very stable nucleosomes at higher temperatures. *J. Mol. Biol.* **245**, 559-567.

2) The compensation I will receive for my study, case preparation, and testimony in this matter is \$100.00 per hour. All travel expenses will be billed at cost.

3) I have not testified as an expert at trial or by deposition within the preceding four years.

- 4) The following includes a complete statement of my opinions to be expressed, the reasons and basis underlying them, the data and other information considered in forming them, and exhibits.

List of Exhibits

- Exhibit 1 — Abstracts from PubMed which contain the word "theory"
- Exhibit 2 — A drawing of the bacterial flagellum from a biochemistry textbook
- Exhibit 3 — Cover page and Table of Contents of the February 6, 1998 issue of the journal *Cell*
- Exhibit 4 — *New York Times* op-ed summarizing the argument for intelligent design
- Exhibit 5 — "Directed Panspermia" by Francis Crick and Leslie Orgel
- Exhibit 6 — *New York Times* article "Biology Text Illustrations More Fiction Than Fact" (sidebar)
- Exhibit 7 — *Philosophy of Science* journal article
- Exhibit 8 — *Biology and Philosophy* journal article
- Exhibit 9 — My chapter from *Debating Design: From Darwin to DNA* (Oxford University Press)
- Exhibit 10 — *Protein Science* journal article
- Exhibit 11 — journal article on "evolutionary potential" by Barry Hall
- Exhibit 12 — *New York Times* op-ed containing my views on teaching the problems of Darwinian theory

1 Intelligent Design as a Scientific Theory

1.1 How the word "theory" is used in the scientific community

1.1.1 "Theory" as a "well-substantiated explanation"

In both common parlance and scientific practice the word "theory" has several meanings¹. The word "theory" is sometimes used in science to indicate, in the words of the National Academy of Sciences, "a well-substantiated explanation of some aspect of the natural world that can incorporate facts, laws, inferences and tested hypotheses."² It is important to remember, however, that even when used in the sense of "well-substantiated" a "theory" is not necessarily either complete or correct. For example, Newton's theory of motion and gravity was superseded by Einstein's theory of relativity.³ Newton's theory had been "well-substantiated" for centuries for bodies of medium size moving at medium speeds (such as cannonballs flying through the air), but did not fit well with bodies moving at very fast velocities, near the speed of light. Another example is the ether theory of the propagation of light, which the eminent 19th century physicist James Clerk Maxwell confidently supported because he thought his equations for electromagnetism required it. The theory proved to be entirely wrong — physicists no longer believe there is such an entity as the "ether".⁴ Thus, as the history of science richly shows, even a "well-substantiated" theory can later be shown to be incomplete or incorrect.

1.1.2 "Theory" as a word that applies to a limited area of science that may be wrong

The word "theory" is also used in scientific practice to apply to proposed explanations that are neither broad in scope nor well-substantiated, which deal with quite limited phenomena that have varying amounts of evidence in their favor, which sometimes are weak or even counterindicated by the data. For example, a search of the online biological database PubMed (which is maintained by the National Library of Medicine, a division of the United States National Institutes of Health) for the word "theory" in abstracts of scientific articles published during or after the year 2000 shows the word is used in varying ways. One article in the *Journal of Theoretical Biology* states that "The membrane pacemaker theory of aging is an extension of the oxidative stress theory of aging."⁵ A paper in the *Journal of Urology* states that "This study does not support the previous theory that urethral sphincter overactivity ... leads to work hyperplasia..."⁶ A paper in the journal *Breast Cancer Research* states their results "do not support the estrogen hypothesis as a unifying theory for the influence of this period."⁷ The point is that the word "theory" is used to indicate a wide range of ideas, some supported by results but limited in scope, some contradicted by results.

Ten abstracts of articles found in the PubMed database are included as Exhibit 1. In each, I have circled the word "theory" in red.

1.1.3 "Theory" applied to each of rival explanations

The word "theory" is also sometimes used in science for each of the opposing explanations which seek to account for the same set of facts. For example, speaking of the continuing

lack of an accepted Darwinian account of the origin of sexual reproduction (which I discuss at more length in section 3.3) an article in the journal *Science* remarked, "Biologists have come up with a profusion of theories since first posing these questions a century ago."⁸ Notice that the word "theories" is used here to indicate competing explanations. Another, more pertinent example is "complexity theory" in biology, which posits that some complex systems can self-organize. Complexity theory has been proposed as an explanation for the origin of such disparate systems as cellular metabolic pathways and the number of different cell types which an organism contains. A leading complexity theorist, Dr. Stuart Kauffman of the University of Calgary, specifically sees aspects of complexity theory as rival explanations to Darwinian theory. In his book *Origins of Order: Self-Organization and Selection in Evolution* (Oxford University Press, 1993) Kauffman wrote:

Darwin and evolutionism stand astride us, whatever the mutterings of creation scientists. But is the view right? Better, is it adequate? I believe it is not. It is not that Darwin is wrong, but that he got hold of only part of the truth. For Darwin's answer to the sources of the order we see all around us is overwhelmingly an appeal to a single singular force: natural selection. It is this single-force view which I believe to be inadequate, for it fails to notice, fails to stress, fails to incorporate the possibility that simple and complex systems exhibit order spontaneously.⁹

Kauffman and other complexity theorists defend their view against both intelligent design theory and Darwinian theory in the recently published book *Debating Design: From Darwin to DNA* (Cambridge University Press, 2004). Thus in scientific usage different "theories" can be rival explanations for the same data.

1.1.4 "Theory" as a singular word applied to a body of multiple, distinct claims

It is critical to realize that in science the word "theory", in the singular, may be applied to a body of multiple, logically-separable claims, some of which may turn out to be true and others false, which can vary widely in the strength of the evidence supporting them and the ease with which they can be tested. In his book *One Long Argument*, in a chapter entitled "Ideological Opposition to Darwin's Five Theories", the eminent, recently-deceased Harvard biologist Ernst Mayr, one of the founders of the "neo-Darwinian synthesis" in the middle of the 20th century, stressed that what is commonly called "Darwin's Theory" actually contains at least five distinct claims.

In both scholarly and popular literature one frequently finds references to "Darwin's theory of evolution," as though it were a unitary entity. In reality, Darwin's "theory" of evolution was a whole bundle of theories, and it is impossible to discuss Darwin's evolutionary thought constructively if one does not distinguish its various components. The current literature can easily leave one perplexed over the disagreements and outright contradictions among Darwin specialists, until one realizes that to a large extent these differences of opinion are due to a failure of some of these students of Darwin to appreciate the complexity of his paradigm.¹⁰

Mayr lists those five separate claims as:

- (1) *Evolution as such.* This is the theory that the world is not constant nor recently created nor perpetually cycling but rather is steadily changing and that organisms are transformed in time.
- (2) *Common descent.* This is the theory that every group of organisms descended from a

- common ancestor and that all groups of organisms, including animals, plants, and microorganisms, ultimately go back to a single origin of life on earth.
- (3) *Multiplication of species.* This theory explains the origin of the enormous organic diversity. It postulates that species multiply, either by splitting into daughter species or by "budding," that is, by the establishment of geographically isolated founder populations that evolve into new species.
 - (4) *Gradualism.* According to this theory, evolutionary change takes place through the gradual change of populations and not by the sudden (saltational) production of new individuals that represent a new type.
 - (5) *Natural selection.* According to this theory, evolutionary change comes about through the abundant production of genetic variation in every generation. The relatively few individuals who survive, owing to a particularly well-adapted combination of inheritable characters, give rise to the next generation.¹¹

The strength of evidence showing that, say, "the world is not constant" (the first claim) is disputed by virtually no one. On the other hand, the strength of evidence showing that natural selection is the sole or principle mechanism of that change in all cases (the fifth claim) is much weaker — necessarily so, since one has to recognize that change has occurred before one attempts to explain what has caused the change. Mayr writes that in the years after Darwin published his theory, although most scientists accepted change in the world and common descent, few scientists thought that the mechanism of natural selection was convincing, as shown in the table below reproduced from Mayr's book.

TABLE 1¹²

The composition of the evolutionary theories of various evolutionists. All these authors accepted a fifth component, that of evolution as opposed to a constant, unchanging world.

	Common descent	Multiplication of species	Gradu- alism	Natural selection
Lamarck	No	No	Yes	No
Darwin	Yes	Yes	Yes	Yes
Haeckel	Yes	?	Yes	In part
Neo- Lamarckians	Yes	Yes	Yes	No
T. H. Huxley	Yes	No	No	(No) ^a
de Vries	Yes	No	No	No
T. H. Morgan	Yes	No	(No) ^a	Unimportant

a. Parentheses indicate ambivalence or contradiction.

In terms of the various usages of the word "theory" discussed in section 1.1, the Darwinian claim that change has occurred on the earth is "well-substantiated". On the other hand, *the altogether separate Darwinian claim that natural selection drove all the major changes on earth, or built all the complex biochemical and cellular systems, is a "theory" in the much weaker sense of a "hypothesis" or "proposed explanation", with much less hard evidence in its favor.*

1.2 How intelligent design theory fits the definition of a scientific theory

1.2.1 The basic claim of intelligent design theory

Intelligent design (ID) theory proposes that the origin of some aspects of living organisms is best explained as the result of deliberate intelligent design, rather than as the result of such unintelligent processes as the self-organization proposed by complexity theory or the natural selection proposed by Darwinian theory. As such, it is crucial to keep in mind that, much like complexity theory, **intelligent design theory focuses exclusively on the proposed mechanism of how complex biological structures arose**. In other words, **ID focuses exclusively on the fifth claim of Darwinism (Natural selection) in Ernst Mayr's list on the preceding page**, and does not concern any of the other claims.

No matter what some people in the general public may say or hope, ID theory does not concern the age of the earth or common descent or any other claims of Darwinian theory except its proposed mechanism. Rather, ID theory exclusively focuses on the question of whether the complex features of organisms are best explained as the result of intelligent or unintelligent causes.

1.2.2 Definition used here for "scientific theory"

Is intelligent design a "scientific theory"? As shown in section 1.1 the word "theory" has multiple meanings. It is my opinion that the relevant dictionary definition for that term in the present context is "the analysis of a set of facts in their relation to one another".¹³ In other words, a "theory" is a proposed explanation for a set of facts. I have argued in the philosophy of science journal *Biology and Philosophy* that a "scientific" theory is a theory which is constructed solely on the foundation of empirical facts about the natural world and logical inferences.¹⁴ Since it is supposed to be based solely on empirical facts and logical inferences, a "scientific" theory should not tailor its claims to agree with authoritative sources, such as the scriptures of any religion or the statements of any religious or governmental leaders, nor should it tailor its claims to disagree with them. Neither should a "scientific" theory deliberately adjust its claims to agree with prevailing expectations among scientists in general of what sorts of phenomena should exist in the universe, nor should it adjust its claims to disagree with them. Rather, a scientific theory should be developed in utter disregard of any factors other than the physical, empirical evidence.

1.2.3 Why ID is a scientific theory

1.2.3.1 The appearance of design in biology

ID is a "theory" because, as discussed above in 1.2.2, it is a proposed explanation for a set of facts. It is a "scientific" theory because, as in 1.2.2, it is based entirely on empirical, observable facts about biology plus logical inferences. The fact that design is indeed based on empirical, observable facts about biology is evident in the writings of some biologists who are not proponents of intelligent design. For example, Francis Crick, the recently deceased Nobel laureate and co-discoverer of the shape of DNA, wrote that, "Biologists must constantly keep in mind that what they see was not designed, but rather evolved".¹⁵ Apparently they must strive so hard to do this because the appearance of design in life is so strong. Brandeis University biologist David DeRosier, a scientist who does research on

the bacterial flagellum, which is a structure that many bacteria use to swim through liquid, remarked in the science journal *Cell* that "More so than other motors, the flagellum resembles a machine designed by a human."¹⁶ (A copy of a drawing of the bacterial flagellum which appears in the university textbook *Biochemistry* by Voet and Voet is included as Exhibit 2.) In 1998 a special edition of *Cell* was devoted to the topic of "Macromolecular Machines" — that is, structures found in the cell which are literally machines made out of molecules. Articles in the journal had titles such as: "The Cell as a Collection of Protein Machines"; "Polymerases and the Replisome: Machines within Machines"; and "Mechanical Devices of the Spliceosome: Motors, Clocks, Springs, and Things". Commentary on the page containing the Table of Contents effused:

Like the machines invented by humans to deal efficiently with the macroscopic world, protein assemblies contain highly coordinated moving parts. Reviewed in this issue of *Cell* are the protein machines that control replication, transcription, splicing ... —the machines that underlie the workings of all living things.¹⁷

In other words the cell — the foundation of life — contains systems that function like sophisticated, designed machinery. A copy of the cover of the issue of *Cell*, as well as a copy of its table of contents, is included as Exhibit 3.

1.2.3.1.1 Richard Dawkins on the appearance of design in biology

That biology exudes the appearance of design is insisted upon by one of the foremost proponents of Darwin's theory, Oxford University biologist Richard Dawkins. On the first page of his book *The Blind Watchmaker* Dawkins bluntly observes, "Biology is the study of complicated things that give the appearance of having been designed for a purpose."¹⁸ As a Darwinist, Dawkins thinks that in reality natural selection accounts for the appearance of design. Nonetheless, he states strongly that the appearance of design in life is overpowering:

Natural selection is the blind watchmaker, blind because it does not see ahead, does not plan consequences, has no purpose in view. Yet the living results of natural selection overwhelmingly impress us with the appearance of design as if by a master watchmaker, impress us with the illusion of design and planning.¹⁹

Dawkins writes that design can easily be recognized from the physical attributes of a system:

We may say that a living body or organ is well designed if it has attributes that an intelligent and knowledgeable engineer might have built into it in order to achieve some sensible purpose, such as flying, swimming, seeing ...²⁰

Dawkins further points out that a system does not have to be perfect to have the marks of design:

It is not necessary to suppose that the design of a body or organ is the best that an engineer could conceive of. ... But any engineer can recognize an object that has been designed, even poorly designed, for a purpose, and he can usually work out what that purpose is just by looking at the structure of the object.²¹

1.2.3.1.2 The lack of rigorous, detailed Darwinian explanations for the appearance of design in biology

Proponents of Darwin's theory such as Richard Dawkins are convinced that natural selection can account for the strong appearance of design in biology. However, like proponents of complexity theory, proponents of intelligent design theory are skeptical of the Darwinian claim, and deny that random mutation and natural selection have been shown to account for some complex aspects of life.

Some scientists who are not proponents of intelligent design freely admit that Darwinian theory has so far been unable to give rigorous, detailed explanations for the complex biochemical machinery discovered in the cell by modern science. For example, in *The Way of the Cell*, published by Oxford University Press in 2001, while considering the claims of intelligent design proponents Colorado State University emeritus microbiologist Franklin M. Harold wrote that:

[W]e must concede that there are presently no detailed Darwinian accounts of the evolution of any biochemical system, only a variety of wishful speculations.²²

When reviewing my book, *Darwin's Black Box: The Biochemical Challenge to Evolution*, which argued that Darwinian explanations have not yet been given for complex biochemical systems, for the science journal *Nature*, University of Chicago evolutionary biologist Jerry Coyne wrote:

There is no doubt that the pathways described by Behe are dauntingly complex, and their evolution will be hard to unravel.... We may forever be unable to envisage the first proto-pathways.²³

The point is that some scientists who are not at all sympathetic to ID nonetheless admit that Darwinian theory has not given detailed, testable explanations for the Darwinian evolution by random mutation and natural selection of complex biochemical systems in the cell. *Thus the path is open to alternative explanations.*

1.2.3.2 Intelligent design reasoning

Intelligent design proponents start from **the same** observable facts as do other scientists such as Richard Dawkins, and notice **the same** strong resemblance of some biological systems, such as the molecular machinery of the cell, to systems we know to be designed. ID advocates also notice that other theories, such as complexity theory or Darwinian natural selection, have not yet given satisfactory accounts, as many scientists freely admit. Thus, ID proponents see a situation where science has discovered that life contains many systems that strongly appear to have been designed and for which no non-design explanation is in hand. ID then advances the plain, straightforward proposal that perhaps the complex structures and molecular machinery of the cell appear designed because they actually were designed by an intelligent agent. I summarized the intelligent design argument in a recent New York Times op-ed, which is included as Exhibit 4.

Although saying that parts of life were actually designed strikes many scientists and other people as unexpected and unsettling, the logic of the design argument is a simple inductive argument (inductive arguments are common and important in science²⁴): Whenever we see functional systems of a certain degree and type of complexity in our everyday world, we have always found them to be designed. Now we have discovered such complex systems in the cell. Since we currently have no other explanation for the origin of such systems, we are justified in extending the induction to the cell, and concluding that there may have been real design involved in its construction.

1.2.3.2.1 An analogy between intelligent design theory and the theory of the Big Bang

The logic of the argument for intelligent design in biology is in some ways similar to that used when the Big Bang theory was first proposed in the earlier 20th century. In the 19th and early 20th century many physicists thought that the universe was eternal and unchanging. Then it was unexpectedly observed that the light from many galaxies was shifted in wavelength toward the red end of the spectrum. This meant that the galaxies were rapidly moving away from the earth and away from each other. Because this is the pattern observed after an explosion in our common experience, by using inductive reasoning the data could be interpreted to be pointing to the aftermath of a huge explosion of the universe itself. This further suggested that the universe was not eternal and unchanging, but may have had a beginning.²⁵

The Big Bang theory struck some scientists as having religious overtones — perhaps the Big Bang was initiated from outside of nature, perhaps it was even a supernatural creation event — and because of this some scientists disliked the theory.²⁶ But the Big Bang theory was not justified by any religious text or dogma. Rather, it was justified by the strong pattern of an explosion suggested by the data. I have written in the philosophy of science journal *Biology and Philosophy* that I think intelligent design theory is similar to the Big Bang theory in the following respect.²⁷ Some people may worry that the theory has religious overtones, but science should ignore any extra-scientific overtones and focus exclusively on the data. Like the Big Bang, intelligent design is justified by the pattern of the physical data.

1.2.3.3 Scientific theories, intelligent design, and falsifiability

Some philosophers of science have argued that one mark of a scientific theory is that it is falsifiable.²⁸ On the other hand, other philosophers of science strongly disagree that falsifiability is a necessary mark of a scientific theory, or that other simple criteria reliably demarcate science from non-science.²⁹ Some scientists have claimed that a theory of intelligent design is unfalsifiable and therefore is not scientific.³⁰ On the other hand, other scientists have actually advanced scientific claims intended to falsify ID, showing that they think ID is indeed falsifiable.³¹ (I have rebutted their claims that ID had been shown to be wrong, arguing instead that the science behind their claims was either mistaken or not pertinent.)³² I have argued in an article in the philosophy of science journal *Biology and Philosophy* that intelligent design theory is falsifiable.³³ The point is that the necessity for

a “scientific” theory to be falsifiable is disputed, but that, in any event, ID is indeed falsifiable.

2 Intelligent Design is not “Creationism”

2.1 Definition of “creationism”

Is ID “creationism”? To answer that question we must first decide what “creationism” is. Some people use the word “creationism” very broadly to indicate any belief that a supernatural being has affected nature in any way. For example, in 1987 John Maddox, then the editor of the prominent science journal *Nature*, wrote an editorial in *Nature* entitled “Down with the Big Bang” that argued that the Big Bang theory was “philosophically unacceptable”, partly because it gave succor to “creationists”:

Creationists and those of similar persuasions seeking support for their opinions have ample justification in the doctrine of the Big Bang. That, they might say, is when (and how) the Universe was created.³⁴

Nonetheless, the common usage of the word “creationism” indicates “the literal belief in the account of creation given in the Book of Genesis”.³⁵ Thus in popular usage a “creationist” is a person who thinks the world is relatively young, on the order of ten thousand years, and that the major categories of organisms were created ex nihilo by a supernatural being, God.

2.2 ID requires none of the presumptions of creationism

Intelligent design theory is not creationism because intelligent design theory does not require belief in any tenet of creationism. As discussed in section 1.2.1, intelligent design theory focuses ***exclusively*** on the question of whether biological systems, such as the molecular machinery found in cells, exhibit features consistent with actual intelligent design. As interesting as they may be, the topics of when such designing occurred, who did the designing, why it was done, how it was done, and so forth, are further, additional questions — beyond the basic question of whether there is design present — for which the scientific evidence is yet insufficient to draw a firm conclusion.

Interestingly, a “creationist” does not at all have to think that biology shows physical, empirical signs of design. A “creationist” can simply believe in creation based on faith in a religious text, private religious experience, or some other source, without consideration of nature at all. Although he was not a “creationist” in our modern sense of the term, the 19th century Englishman Cardinal John Henry Newman exemplified that attitude. In his *Letters and Diaries* he wrote, “I believe in design because I believe in God, not in a God because I see design.”³⁶ In other words, strong religious faith does not require that biology show any physical evidence of design of the kind of which the Darwinian evolutionary biologist Richard Dawkins writes (see section 1.2.3.1.1), or that Brandeis University biologist David DeRosier sees in the bacterial flagellum (sections 1.2.3.1).

One can be a "creationist" in the popular sense of that word and think biology does show signs of design, but one certainly need not be. For example, as I discussed in my book *Darwin's Black Box*³⁷ and in an article in the philosophy of science journal *Biology and Philosophy*,³⁸ one is free to suppose, based on nonscientific reasons, that the designer was a natural being, such as a time traveler or space alien. (Francis Crick, the Nobel laureate, once proposed in an article entitled "Directed Panspermia" in the science journal *Icarus* that the origin of life on earth may have been the result of the deliberate seeding of life here by spores sent by intelligent space aliens.³⁹ A copy of Crick's article is included as Exhibit 5.) Indeed, there is at least one group (the "Raelians") who profess to believe that humans were designed by space aliens.⁴⁰

One can also hold that the designer is some other, yet unexplained, natural entity. If one decides, on the basis of nonscientific reasons, that the designer is a supernatural being, then one can hold that the designer is a subordinate supernatural being such as the "demiurge" of Plato.⁴¹ If one supposes that the designer is a supreme supernatural being, God, then it may be the God of any religion, such as that of Christianity, Islam, Hinduism, Native American religions, or others. Or one can simply keep an open mind, and think that the question of the identity of the designer has yet to be resolved with any firmness.

One can also think, unlike a "creationist", that a designer set up the universe to unfold in a planned way over immense times, to give rise to the complex structures science has discovered in life, without any discernible exception to natural laws. I discussed this view in a letter to the editor published in the July 2001 edition of the NCSE Reports. The newsletter is published by the National Center for Science Education, which vigorously promotes the teaching of Darwinian evolution in schools. The letter is reproduced below:

In their article "Of Mousetraps and Men: Behe on Biochemistry" (Reports of the NCSE 20, 25-30, 2000), which has just come to my attention, Shanks and Joplin appear to mistakenly attribute to me the contention that irreducibly complex biochemical systems must have been created ex nihilo. I have never claimed that. I have no reason to think that a designer could not have used suitably modified pre-existent material. My argument in *Darwin's Black Box* is directed merely toward the conclusion of design. How the design was effected is a separate and much more difficult question to address. Although creation ex nihilo is a formal possibility, design might have been produced by some other means which involved no discontinuities in natural law, even if the designer is a supernatural being. One possibility is directed mutations. As noted by Brown University biologist Kenneth Miller in *Finding Darwin's God*, "The indeterminate nature of quantum events would allow a clever and subtle God to influence events in ways that are profound, but scientifically undetectable to us. Those events could include the appearance of mutations...." I have no reason to object to that as a route to irreducibly complex systems. I would just note further that such a process amounts to intelligent design, and that while we may be unable to discern the means by which the design is effected, the resultant design itself may be detected in the structure of the irreducibly complex system.

The core claim of intelligent design theory is quite limited. It says nothing directly about how biological design was produced, who the designer was, common descent, or other such questions. Those can be argued separately. It says only that design can be empirically detected in observable features of physical systems. As an important corollary, it also predicts that mindless processes—such as natural selection or the self-organization scenarios favored by Shanks and Joplin—will not be demonstrated to be able to produce irreducible systems of the complexity found in cells.⁴²

None of the possibilities discussed above requires that the picture of the universe developed by modern science be repudiated. The only assertion that intelligent design theory itself properly makes is that some aspects of biology are indeed the product of intelligent design. As discussed earlier in section 1.2.3, this assertion is actually quite consistent with the evidence of biology, although at odds with the claims of Darwinism. The point is that intelligent design theory does not require a person to adhere to any tenet usually associated with the word "creationism". A "creationist" does not have to believe in physically-discernible intelligent design, and an ID proponent does not have to believe in "creationism".

3 What are the gaps and problems with the Darwinian theory of evolution?

3.1 The problem of the origin of new, complex biological features

It is my scientific opinion that the primary problem with Darwin's theory of evolution is the lack of detailed, testable, rigorous explanations for the origin of new, complex, biological features, as explained above in section 1.2.3.1.2. This problem was recognized in the 19th century, shortly after Darwin published *The Origin of Species*, by biologists such as St. George Mivart ("What is to be brought forward [against Darwin's theory] may be summed up as follows: That "Natural Selection" is incompetent to account for the incipient stages of useful structures..."⁴³, and continues to be a problem today. Although vague stories and speculations are sometimes offered, rarely are such stories testable in a way that could falsify the claim that the complex feature was produced in a Darwinian fashion. For example, as stated above in section 1.2.3.1.2, in the case of the molecular machinery found in cells Franklin M. Harold wrote that: "[W]e must concede that there are presently no detailed Darwinian accounts of the evolution of any biochemical system, only a variety of wishful speculations." And Jerry Coyne wrote "There is no doubt that the pathways described by Behe are dauntingly complex, and their evolution will be hard to unravel.... We may forever be unable to envisage the first proto-pathways." *It is extremely difficult or impossible to test — or even meaningfully critique — "wishful speculations" or un-envisaged proto-pathways.*

It should be strongly emphasized that under this broad category of difficulties lies much of the structure and development of life, including: the existence of the genetic code; transcription of DNA; translation of mRNA; the structure and function of the ribosome; the structure of the cytoskeleton; nucleosome structure; the development of new protein-protein interactions; the existence of the proteosome; the existence of the endoplasmic reticulum; the existence of motility organelles such as the bacterial flagellum and the eukaryotic cilium; the development of the pathways for the construction of the cilium and flagellum; the existence of the defensive apparatus such as the immune system and blood clotting system; and much else. The existence of such unresolved difficulties for Darwinian theory at the molecular level of life makes it reasonable to wonder if a Darwinian framework is the right way to approach such questions. It also makes it reasonable to wonder if Darwinian processes explain major new features of life at higher levels, such as the level of organs and organisms.

3.2 The problem of falsification

There are other major difficulties and problems for Darwin's theory as well. One is the great difficulty in falsifying it. That is, in finding a fact of nature that would be taken by Darwinists as evidence against their theory. For example, for many years in biology textbooks students were shown drawings of vertebrate embryos that looked remarkably similar. The embryos were drawn by the 19th century embryologist Ernst Haeckel, an admirer of Darwin. The striking similarity was thought to strongly support Darwin's theory, that the different classes of vertebrates descended by natural selection from a common ancestor. The rationale for thinking so was given in the widely-used, college-level textbook *Molecular Biology of the Cell*, where president of the National Academy of Sciences Bruce Alberts and other co-authors wrote that:

Early developmental stages of animals whose adult forms appear radically different are often surprisingly similar... Such observations are not difficult to understand.... The early cells of an embryo are like cards at the bottom of a house of cards—a great deal depends on them, and even small changes in their properties are likely to result in disaster.⁴⁴

In other words, evolution would be expected to conserve the structure of the early embryos, inherited from a common ancestor. Natural selection would not be expected to change such a "locked-in", fundamental structure.

However, in 1997 an international team led by the British embryologist Michael Richardson showed that Haeckel's drawing were very misleading, and that there were significant differences between the embryos. A story entitled "Haeckel's embryos: fraud rediscovered" in the journal *Science* put it this way:

Not only did Haeckel add or omit features ... but he also fudges the scale to exaggerate similarities among species, even when there were 10-fold differences in size. Haeckel further blurred differences by neglecting to name the species in most cases, as if one representative was accurate for an entire group of animals. In reality ... even closely related embryos such as those of fish vary quite a bit.⁴⁵

Nonetheless, the discovery that the embryos looked very different from what they were pictured in textbooks did not at all cause Bruce Alberts or other scientists to question Darwinian theory. Yet if a theory is equally compatible with one result (nearly identical embryos) and its opposite (variable embryos) than how can it be rigorously tested? If Darwinian theory is compatible with false data, such as the original drawings of Haeckel, then how can we know if the theory is wrong? A story from the *New York Times*, "Biology Text Illustrations More Fiction Than Fact" (which is a sidebar in the longer story "Darwin vs. Design: Evolutionists' New Battle"), concerning the case of Haeckel's embryos is included as Exhibit 6.⁴⁶

3.3 The "crisis" of sex

Another major, longstanding difficulty of Darwinian theory that is little appreciated by the general public is the problem of sexual reproduction. It was recognized as early as the late

19th century that Darwinian theory predicted sexual reproduction should be rare. The reason is that in sexual reproduction a given organism only gets half of its genes reproduced in any particular one of its offspring; its sexual partner contributes the other half. However, if an organism reproduced asexually, then it could contribute all of its genes to each of its offspring. The reason that, contrary to the straightforward expectation of Darwinian theory, sex is common has been debated for over a hundred years. There are so many competing ideas about why sexual reproduction should occur that A. S. Kondrashov, in an article in the *Journal of Heredity* in 1993, found it necessary to try to classify all the hypotheses into groups to keep better track of them!

After more than a century of debate, the major factors of the evolution of reproduction are still obscure. During the past 25 years, hypotheses have become so numerous and diverse that their classification is a necessity. The time is probably ripe for this: no fundamentally new hypothesis has appeared in the last 5 years, and I would be surprised—and delighted—if some important idea remains unpublished.⁴⁷

The debate continues, with various theories offered by various scientists, but with no accepted resolution of the problem. For example in 1998 the journal *Science* devoted a special section of one issue to the evolution of sex, with articles such as “Why Sex? Putting Theory to the Test” and “Why Sex and Recombination?” One introductory article remarked,

Yet how sex began and why it thrived remain a mystery. ... Why did sex overtake asexual reproduction some billion or more years ago, and why does it continue to upstage asexuals? ... Biologists have come up with a profusion of theories since first posing these questions a century ago. ... Sex is a paradox in part because if nature puts a premium on genetic fidelity, asexual reproduction should come out ahead. It transmits, intact, a single parental genome that is by definition successful. Sexual reproduction, on the other hand, involves extensive makeovers of the genome. The production of gametes requires recombination, in which the two copies of each chromosome pair up and exchange DNA. Fertilization, in which genes from different parents fuse, creates yet more genetic combinations. All this shuffling is more likely to break up combinations of good genes than to create them—yet nature keeps reshuffling the deck.⁴⁸

(Notice again that the word “theories” is used above to mean competing, tentative ideas — not “well-substantiated explanations”.) In his 1975 book *Sex and Evolution* the prominent evolutionary biologist George C. Williams wrote:

This book is written from a conviction that the prevalence of sexual reproduction in higher plants and animals is inconsistent with current evolutionary theory ... there is a kind of crisis at hand in evolutionary biology ...⁴⁹

While discussing Williams remarks on sex, in 2004 Richard Dawkins wrote:

Maynard Smith and Hamilton said similar things. It is to resolve this crisis that all three Darwinian heroes, along with others of the rising generation, laboured. I shall not attempt an account of their efforts, and certainly I have no rival solution to offer myself.⁵⁰

Yet if Darwinian theory has no good account for sexual reproduction, then the very heart of the theory of evolution — the differential reproduction of organisms — is floating in midair.

A theory of evolution that predicts most species should be asexual is like a theory of gravity

that predicts that most objects will fall up. Either conundrum should make a reasonable person wonder if the proposed theory might be missing some large piece of the puzzle. Furthermore, if Darwinists have tolerated such a large, acknowledged difficulty in the center of their theory for over a hundred years, then one might wonder if they are unreasonably attached to it. Students should be aware of this.

It should be emphasized that the problem with sex is not simply the problem of how the intricate machinery of meiosis, recombination, and other sexual processes could develop in the gradual, undirected manner that Darwinian theory envisions. The theory encounters that difficulty in the explanation of *all* complex biological systems. Rather, the difficulty in explaining sex is the question of why, on Darwinian principles, it should exist at all, even if there were a gradual way to develop it. On straightforward Darwinian principles, sex appears detrimental to the interests of the organism.

4 The origin of life

A major problem for Darwin's theory is the unsolved mystery of the origin of life. Darwin's theory doesn't itself deal with the origin of life; rather, it presupposes that life was present on earth in a form that would be able to undergo evolution by random mutation and natural selection. Even in principle, Darwin's theory cannot account for the origin of life, because the theory concerns the reproduction of already-living organisms. Thus before the beginning of life the earth was missing a prerequisite ingredient for Darwinian evolution to occur.

The problem that the origin of life poses for Darwin's theory is the following. If the beginning of life required something extra, something in addition to the unintelligent operation of natural processes that Darwin's theory invokes, then it would be fair for a curious inquirer to wonder if those other processes ended with the beginning of life, or if they continued to operate throughout the history of life. The acknowledgment of difficulties with the origin of life would likely make it more urgent that Darwinists actually demonstrate that random mutation and natural selection can do what they claim, rather than relying on the presumption that they can.

The importance of the origin of life to Darwin's theory is seen in the fact that high school biology textbooks include a section dealing with the topic. This often leads into discussion of the first cells and then into Darwinian evolution, so that it can appear to the student to be a seamless process. Sometimes a text gives students little warning (or the warning is not emphasized so that students easily overlook it) that the origin of life is an unsolved problem that has remained a mystery despite fifty years of active scientific investigation.

It is easy to find comments by knowledgeable scientists that attest to the lack of progress in the field of origin of life studies. For example, in a recent interview with the PBS science program Nova, the distinguished paleontologist Andrew Knoll, who is the Fisher Professor of Natural History at Harvard University and a leading expert on early life on earth, remarked: "The short answer is we don't really know how life originated on this planet." In response to the interviewer's question, "Will we ever solve the problem [of the origin of

life]?", Knoll responded, "I imagine my grandchildren will still be sitting around saying that it's a great mystery..."⁵¹

The widely-used, university-level textbook "Biochemistry" by Voet and Voet introduces a section on the origin of life with the following remarks:

In the remainder of this section, we describe the most widely favored scenario for the origin of life. *Keep in mind, however, that there are valid scientific objections to this scenario as well as to the several others that have been seriously entertained, so that we are far from certain as to how life arose.*⁵² [italics in the original]

In other words, like with the problem of sex, there is a profusion of theories, none of which is satisfactory.

In its booklet *Science and Creationism* the National Academy of Sciences called the problem of the origin of life "seemingly intractable." Nonetheless, the National Academy writes:

For those who are studying the origin of life, the question is no longer whether life could have originated by chemical processes involving nonbiological components. The question instead has become which of many pathways might have been followed to produce the first cells.⁵³

This statement subtly shifts the spotlight away from the actual scientific *problem* of the origin of life and onto the subjective *attitudes* of workers in the field. In effect it encourages teachers (to whom the booklet is addressed) to inculcate in their students the presumption that the problem of the origin of life must be addressed in the framework of unintelligent "chemical processes involving nonbiological components". This despite the fact that such a framework has been unsuccessful over the course of half of a century. Students are not encouraged to think, or given any reason to think, that such a framework might possibly be wrong. Students are encouraged to follow in the footsteps of the failures of the past fifty years.

One can also discern in another, quite remarkable passage in *Science and Creationism* the desire to inculcate into students the presumption that "chemical processes involving nonbiological components" simply must be responsible for the origin of life. There the National Academy of Sciences speaks glowingly of a particular *theological* stance, called "theistic evolution", as if the Academy — an organization chartered by the federal government — were expert on religious matters.

Many religious persons, including many scientists, hold that God created the universe and the various processes driving physical and biological evolution and that these processes then resulted in the creation of galaxies, our solar system, and life on Earth. This belief, which sometimes is termed "theistic evolution," is not in disagreement with scientific explanations of evolution. Indeed, it reflects the remarkable and inspiring character of the physical universe revealed by cosmology, paleontology, molecular biology, and many other scientific disciplines.⁵⁴

A teacher reading that section could easily pick up the Academy's apparent attitude toward religion: theistic evolution, where laws operate continuously — good; religious ideas

requiring interruption of natural laws — bad. A teacher influenced by the Academy's booklet might possibly attempt to influence the religious beliefs of students in the same way.

5 The scientific controversy over intelligent design

My book, *Darwin's Black Box: The Biochemical Challenge to Evolution*, presented the argument that Darwinian processes are unlikely explanations for the biochemical complexity that modern science has found in the cell. Instead, the book argued, a more likely explanation is deliberate intelligent design. Since shortly after the book was published in 1996 scientists who support Darwin's theory of evolution by random mutation and natural selection have offered arguments to try to refute the contention of intelligent design. In turn I have offered counterarguments to show why the Darwinian arguments fail. I think it is safe to say that so far neither side has been persuaded by the other's arguments. Below I will list some of the articles that have been published on both sides.

1) In a symposium published by *Boston Review* in its Feb/March 1997 issue, a dozen academics traded essays arguing the relative merits of intelligent design, Darwinism, and other ideas for explaining the development of life. The essays are available on line.⁵⁵ Contributing authors include:⁵⁶

Michael Behe, professor of biological sciences, Lehigh University
 Phillip E. Johnson, professor of law, University of California, Berkeley,
 David Berlinski, a writer and mathematician,
 Jerry A. Coyne, professor of evolutionary biology, University of Chicago
 Russell F. Doolittle, professor of biochemistry, University of California, San Diego
 Douglas J. Futuyma, professor of evolutionary biology, State University of New York, Stony Brook
 Robert DiSilvestro, professor of nutritional biochemistry, Ohio State University
 Michael Ruse, professor of philosophy, University of Guelph
 James A. Shapiro, professor of biochemistry, University of Chicago
 Daniel Dennett, professor of philosophy, Tufts University
 H. Allen Orr, professor of evolutionary biology, University of Rochester

2) In 1999 Kenneth Miller, a professor of biology at Brown University, published *Finding Darwin's God*⁵⁷ (HarperCollins). In the book he defended Darwinian evolution. One of the chapters of the book, Chapter 5 "God the Mechanic", criticizes my argument in *Darwin's Black Box* for intelligent design, and offers scientific arguments against it.

3) In 2001 Robert T. Pennock, professor of philosophy at Michigan State University, edited a book entitled *Intelligent Design Creationism and Its Critics*⁵⁸, which was published by MIT Press. The book collected dozens of essays. Each essay by a proponent of intelligent design was subjected to several critical essays by opponents. Proponents usually were not given space to respond to criticisms. Several of the essays concerned the scientific claims of intelligent design.

- 4) In 1999 Shanks and Joplin published a review of *Darwin's Black Box* in the journal *Philosophy of Science*⁵⁹. They argued that the idea of irreducible complexity (which I discussed in the book) was incorrect, and that complex biochemical systems could develop through a means they called "redundant complexity."
- 5) In 2000 I replied to Shanks and Joplin's criticisms in an article also published in *Philosophy of Science*⁶⁰. I argued that their criticisms of irreducible complexity were themselves flawed. A copy of the article is included as Exhibit 7.
- 6) In 2001 I published an article entitled "Reply to my critics: A response to reviews of *Darwin's Black Box: the biochemical challenge to evolution*" in the journal *Biology and Philosophy*⁶¹. The article responds to many of the criticisms in books and articles listed above, including those of scientists Kenneth Miller, Russell Doolittle, H. Allen Orr, Jerry Coyne. A copy of the article is included as Exhibit 8.
- 7) In 2000 Thornhill and Ussery published an article in the *Journal of Theoretical Biology* arguing against the concept of irreducible complexity.⁶²
- 8) In 2000 a conference organized by William Dembski was held at Baylor University. It was entitled "the Nature of Nature", and brought together intelligent design proponents and opponents, including many scientists, mathematicians, and philosophers, as well as several Nobel laureates and members of the National Academy of Sciences.⁶³
- 9) In 2003 Lenski et al published an article in the journal *Nature* entitled "The evolutionary origin of complex features".⁶⁴ The article concerned the ability of a computer program to develop the ability to perform new functions. It was intended to be a model for how biological features might develop in organisms and possibly get around the difficulty of irreducible complexity.
- 10) In 2004 Young and Edis edited a volume entitled *Why Intelligent Design Fails: A Scientific Critique of the New Creationism*⁶⁵ (Rutgers University Press) which, as its title suggests, offered scientific arguments against intelligent design.
- 11) In 2004 Dembski and Ruse edited *Debating Design: From Darwin to DNA*⁶⁶ (Cambridge University Press), which included contributions from proponents and opponents of intelligent design, as well as contributions from complexity theorists, who disagree with some of the claims of both intelligent design and Darwinian theory, as well as theistic evolutionists. A copy of my chapter in the volume, which responds to criticisms of irreducible complexity and intelligent design, is included as Exhibit 9.
- 12) In 2004 Behe and Snoke published an article in the journal *Protein Science* entitled "Simulating evolution by gene duplication of protein features that require multiple amino acid residues"⁶⁷. The article attempts to show the difficulty of evolving a new protein feature by random mutation and natural selection when multiple changes are needed for a new function. A copy of the article is included as Exhibit 10.

One point of this compilation is to show that some Darwinian scientists have responded to

intelligent design with scientific arguments that attempt to falsify it — to show ID to be incorrect. Although I think their scientific arguments are incorrect, the fact that scientists offer such arguments demonstrates that intelligent design is amenable to scientific investigation and criticism. It is therefore a scientific claim.

6 The utility of design as a scientific theory

6.1 A scientific theory does not need to be utilitarian

A scientific theory does not have to have an immediately-obvious utilitarian application to be correct. One of the purposes of science is simply to describe nature accurately. If a theory does that, or at least is better than competing theories at describing nature, then it is fulfilling an important purpose of a scientific theory. In the view of proponents of intelligent design theory, ID more accurately describes what we observe in nature than do competing theories. As explained earlier, some scientists admit that Darwinian theory does not have detailed, rigorous explanations for some of the complex systems that have been discovered in the cell. And some scientists such as Richard Dawkins readily admit that aspects of biology strongly appear to have been designed. Thus it is reasonable to conclude, as ID proponents do, that intelligent design is a more accurate description of aspects of nature than other theories.

6.2 Where is the border between design and unintelligent natural processes?

One use of a theory of intelligent design might be to prod scientists to look for limits to the efficacy of the Darwinian processes of random mutation and natural selection, which might lead to describing nature more accurately. If one has reason to believe, as proponents of ID do, that not all of biology can be explained by natural selection, then one can begin to look for the borders of Darwinian processes. A question such as, what are the limits of Darwinian processes in explaining life on earth?, does not easily occur to a Darwinist, who takes it as an assumption that Darwinian processes explain most complex biological systems. A small step toward addressing such questions was recently taken by myself and David Snoke, a professor of physics at the University of Pittsburgh. We recently published an article in the journal *Protein Science* entitled "Simulating evolution by gene duplication of protein features that require multiple amino acid residues". The article attempts to show the difficulty of evolving a new protein feature by random mutation and natural selection when multiple changes are needed for a new function. A copy of the article is included as Exhibit 10.

6.3 Health implications for a limit to Darwinian evolution

Although a scientific theory does not have to have practical implications in order to be correct, a theory of intelligent design nonetheless might be important in understanding such things as the limits of the development of antibiotic resistance, which of course could have great importance for public health. Here is why. It is well known that bacteria and other microorganisms can develop resistance to some antibiotic drugs, and that this is a

formidable public health threat. It is less well known that some bacteria have been unable to develop resistance to some drugs. The reason is that some antibiotic resistance genes do not have the "evolutionary potential" to develop resistance. This is exemplified in some recent articles from the laboratory of Professor Barry G. Hall at the University of Rochester. Although he is not an advocate of intelligent design, Professor Hall nonetheless does not automatically assume that Darwinian processes can do everything. For example, he writes in a paper in the journal *Antimicrobial Agents and Chemotherapy*: "Instead of assuming that metallo- β -lactamases will evolve rapidly, it would be highly desirable to accurately predict their evolution in response to carbapenem selection." Using a method he developed, he predicts that bacteria will be unable to develop resistance to an antibiotic called imipenem. He writes in the abstract of his article: "The results predict, with >99.9% confidence, that even under intense selection the IMP-1 β -lactamase will not evolve to confer increased resistance to imipenem." A copy of Hall's article is included as Exhibit 11.

If intelligent design theory is correct, and there are limits to what unaided nature can do, then if we understand in more detail what those limits are, we may be able to design more effective antibiotics, ones to which bacteria will be unable to develop resistance.

I should emphasize that this is just one possible application of intelligent design theory, which may or may not be easily successful. The overarching point, however, is that approaching the study of biology from an intelligent design perspective may afford insights that do not come easily to workers who have a Darwinian perspective.

7 The age-appropriateness of discussing difficulties with Darwin's theory in 9th grade

It is my opinion that discussing difficulties with Darwin's theory is quite appropriate for students in 9th grade, or in whichever grade a high school biology course is given. The reason is that the problems with Darwin's theory are no more difficult to understand than the advantages of Darwin's theory. Indeed, the difficulties are often just the reverse of the advantages. Even molecular difficulties with the theory are appropriate for high school students. In many high school biology texts, students are taught of the underlying chemical, biochemical, genetic, and cellular bases of life. If the students can understand such topics, then they can understand difficulties that arise for Darwinian theory at this level. My opinions on this topic are summarized in a *New York Times* op-ed piece "Teach Evolution—And Ask Hard Questions", which is included as Exhibit 12.

Signed: Michael J. Behr Date: 3/24/05

Endnotes

1. Merriam-Webster, I. 1986. *Webster's ninth new collegiate dictionary*. Merriam-Webster: Springfield, Mass., U.S.A.
2. National Academy of Sciences. 1999. *Science and creationism: a view from the National Academy of Sciences*, 2nd ed. ed. National Academy Press: Washington, DC, p. 2.
3. "relativistic mechanics." Encyclopædia Britannica from Encyclopædia Britannica Online. <<http://search.eb.com/eb/article?tocId=9110309>> [Accessed March 15, 2005].
4. "ether." Encyclopædia Britannica from Encyclopædia Britannica Online. <<http://search.eb.com/eb/article?tocId=9033121>> [Accessed March 15, 2005].
5. Hulbert, A.J. 2005. On the importance of fatty acid composition of membranes for aging. *J. Theor. Biol.* **234**:277-288.
6. Andrich, D.E., Rickards, D., Landon, D.N., Fowler, C.J., and Mundy, A.R. 2005. Structural assessment of the urethral sphincter in women with urinary retention. *J. Urol.* **173**:1246-1251.
7. Hodgson, M.E., Newman, B., and Millikan, R.C. 2004. Birthweight, parental age, birth order and breast cancer risk in African-American and white women: a population-based case-control study. *Breast Cancer Res.* **6**:R656-R667.
8. Wuethrich, B. 1998. Why sex? Putting theory to the test. *Science* **281**:1980-1982.
9. Kauffman, S.A. 1993. *The origins of order: self-organization and selection in evolution*. Oxford University Press: New York, p. xiii.
10. Mayr, E. 1991. *One long argument: Charles Darwin and the genesis of modern evolutionary thought*. Harvard University Press: Cambridge, Mass., p. 35.
11. Mayr, 1991, pp. 36-37.
12. Mayr, 1991, p. 37.
13. Merriam-Webster, I. 1986. *Webster's ninth new collegiate dictionary*. Merriam-Webster: Springfield, Mass., U.S.A.
14. Behe, M.J. 2001. Reply to my critics: A response to reviews of *Darwin's Black Box: the biochemical challenge to evolution*. *Biology and Philosophy* **16**:685-709.
15. Crick, F. 1988. *What mad pursuit: a personal view of scientific discovery*. Basic Books: New York, pp. 138-139.

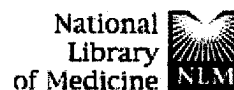
16. DeRosier,D.J. 1998. The turn of the screw: the bacterial flagellar motor. *Cell* **93**:17-20.
17. *Cell*, "Review Issue on Macromolecular Machines", February 6, 1998,
18. Dawkins,R. 1986. *The blind watchmaker*. Norton: New York, p. 1.
19. Dawkins, 1986, p. 21.
20. Dawkins, 1986, p. 21.
21. Dawkins, 1986, p. 21.
22. Harold,F.M. 2001. *The Way of the Cell*. Oxford University Press: Oxford, p. 205.
23. Coyne,J.A. 1996. God in the details. *Nature* **383**:227-228.
24. "logic." Britannica Student Encyclopedia from Encyclopædia Britannica Online. <<http://search.eb.com/ebi/article?tocId=204014>> [Accessed March 18, 2005].
25. "Lemaître, Georges." Encyclopædia Britannica from Encyclopædia Britannica Online. <<http://search.eb.com/eb/article?tocId=9047718>> [Accessed March 16, 2005].
26. Maddox,J. 1989. Down with the Big Bang. *Nature* **340**:425.
27. Behe,M.J. 2001. Reply to my critics: A response to reviews of *Darwin's Black Box: the biochemical challenge to evolution*. *Biology and Philosophy* **16**:685-709.
28. Thornton, Stephen, "Karl Popper", The Stanford Encyclopedia of Philosophy (Winter 2002 Edition), Edward N. Zalta (ed.), URL = <<http://plato.stanford.edu/archives/win2002/entries/popper/>>; Ruse M., ed., 1996, "But is it Science?: The Philosophical Question in the Creation/Evolution Controversy," Prometheus Books: Amherst NY.
29. Laudan L., 1996, "Science at the Bar—Causes for Concern," in Ruse, 1996.
30. see my discussion of this in Behe, 2001, pp. 696-697.
31. Shanks,N. and Joplin,K.H. 1999. Redundant complexity: A critical analysis of intelligent design in biochemistry. *Phil. Sci.* **66**:268-282; Doolittle,R.F. A delicate balance. *Boston Review* [Feb/March], 28-29. 1997; Miller,K.R. 1999. *Finding Darwin's God: a scientist's search for common ground between God and evolution*. Cliff Street Books: New York.
32. Behe,M.J. 2000. Self-organization and irreducibly complex systems: A reply to Shanks and Joplin. *Phil. Sci.* **67**:155-162; Behe,M.J. 2001. Reply to my critics: A response to reviews of *Darwin's Black Box: the biochemical challenge to evolution*.

- Biology and Philosophy* **16**:685-709; Behe,M.J. 2004. Irreducible Complexity: Obstacle to Darwinian Evolution. In *Debating Design: from Darwin to DNA*. (eds. WA Dembski and M Ruse), pp 352-370. Cambridge University Press: Cambridge.
33. Behe, 2001.
34. Maddox,J. 1989. Down with the Big Bang. *Nature* **340**:425.
35. creationism." TheFreeDictionary.com. <<http://www.thefreedictionary.com/creationism>> [Accessed March 15, 2005].
36. www.newmanreader.org/biography/ward/volume2/chapter28.html
37. Behe,M.J. 1996. *Darwin's black box: the biochemical challenge to evolution*. The Free Press: New York, Ch. 11.
38. Behe, 2001.
39. Crick,F. and Orgel,L.E. 1973. Directed panspermia. *Icarus* **19**:341-346.
40. www.rael.org
41. "Demiurge." Encyclopædia Britannica from Encyclopædia Britannica Online. <<http://search.eb.com/eb/article?tocId=9029894>> [Accessed March 16, 2005].
42. Behe,M.J. 2001. Letter to the editor, "Behe replies to Shanks and Joplin",*NCSE Reports* **21**: 4.
43. Mivart,S. 1871. *On the genesis of species*, 2nd ed ed. Macmillan: London, p. 21.
44. Alberts,B. 1994. *Molecular biology of the cell*, 3rd ed. ed. Garland Pub.: New York.
45. Pennisi,E. 1997. Haeckel's embryos: fraud rediscovered. *Science* **277**:1435.
46. Glanz,J. Biology Text Illustrations More Fiction Than Fact. New York Times , A18. 4-8-2001.
47. Kondrashov,A.S. 1993. Classification of hypotheses on the advantage of ammixis. *J. Hered.* **84**:372-387.
48. Wuethrich,B. 1998. Why sex? Putting theory to the test. *Science* **281**:1980-1982.
49. Quoted in Dawkins,R. 2004. *The ancestor's tale: a pilgrimage to the dawn of evolution*. Houghton Mifflin: Boston, p. 424.
50. Dawkins, 2004, p. 424.
51. www.pbs.org/wgbh/nova/origins/knoll.html

52. Voet,D. and Voet,J.G. 2004. *Biochemistry*, 3rd ed ed. J. Wiley & Sons: New York, p. 31.
53. National Academy of Sciences, 1999, p. 6.
54. National Academy of Sciences 1999, p. 7.
55. www.bostonreview.net/evolution.html
56. The institutional affiliations are those at the time the essays were published. Several authors have since switched jobs.
57. Miller, 1999.
58. Pennock,R.T. 2001. *Intelligent design creationism and its critics: philosophical, theological, and scientific perspectives*. MIT Press: Cambridge, Mass.
59. Shanks,N. and Joplin,K.H. 1999. Redundant complexity: A critical analysis of intelligent design in biochemistry. *Phil. Sci.* **66**:268-282.
60. Behe, 2000.
61. Behe, 2001.
62. Thornhill,R.H. and Ussery,D.W. 2000. A classification of possible routes of Darwinian evolution. *J. Theor. Biol.* **203**:111-116.
63. www.designinference.com/documents/2000.04.nature_of_nature.htm
64. Lenski,R.E., Ofria,C., Pennock,R.T., and Adami,C. 2003. The evolutionary origin of complex features. *Nature* **423**:139-144.
65. Young,M. and Edis,T. 2004. *Why intelligent design fails: a scientific critique of the new creationism*. Rutgers University Press: New Brunswick, N.J.
66. Dembski,W.A. and Ruse,M. 2004. *Debating design: from Darwin to DNA*. Cambridge University Press: Cambridge, U.K.
67. Behe,M.J. and Snoke,D.W. 2004. Simulating evolution by gene duplication of protein features that require multiple amino acid residues. *Protein Sci.* **13**:2651-2664.

Exhibit 1

Abstracts from PubMed which
contain the word "theory"



My NCBI

[\[Sign In\]](#) [\[Register\]](#)[All Databases](#)[PubMed](#)[Nucleotide](#)[Protein](#)[Genome](#)[Structure](#)[OMIM](#)[PMC](#)[Journals](#)[Books](#)

Search

PubMed

for

Go

Clear

☒ Limits[Preview/Index](#)[History](#)[Clipboard](#)[Details](#)[About Entrez](#)[Text Version](#)[Entrez PubMed](#)[Overview](#)[Help | FAQ](#)[Tutorial](#)[New/Noteworthy](#)[E-Utilities](#)[PubMed Services](#)[Journals Database](#)[MeSH Database](#)[Single Citation Matcher](#)[Batch Citation Matcher](#)[Clinical Queries](#)[LinkOut](#)[My NCBI \(Cubby\)](#)[Related Resources](#)[Order Documents](#)[NLM Catalog](#)[NLM Gateway](#)[TOXNET](#)[Consumer Health](#)[Clinical Alerts](#)[ClinicalTrials.gov](#)[PubMed Central](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000, English**

Display

Abstract

Show: 20

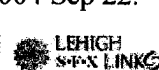
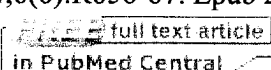
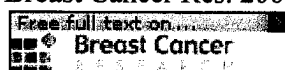
Sort

Send to

Text

All: 1

Review: 0

☐ 1: Breast Cancer Res. 2004;6(6):R656-67. Epub 2004 Sep 22.[Related Articles, Links](#)**Birthweight, parental age, birth order and breast cancer risk in African-American and white women: a population-based case-control study.****Hodgson ME, Newman B, Millikan RC.**

Department of Epidemiology, School of Public Health, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina, USA.

ehodgson@email.unc.edu

INTRODUCTION: Much recent work has focused on hypotheses that very early life exposures influence adult cancer risk. For breast cancer it has been hypothesized that high in utero estrogen exposure may increase risk. **METHODS:** We used data from the Carolina Breast Cancer Study, a population-based case-control study of incident breast cancer in North Carolina, to examine associations for three possible surrogates of high prenatal estrogen exposure: weight at birth, maternal age, and birth order. We also examined paternal age. Birthweight analyses were conducted for white and African-American women born in North Carolina on or after 1949 (196 cases, 167 controls). Maternal age was analyzed for US born participants younger than 49 years of age (280 cases, 236 controls). **RESULTS:** There was a weak inverse association between birthweight in the highest tertile and breast cancer overall (odds ratio [OR] 0.7, 95% confidence interval [CI] 0.4-1.2), although associations differed by race (OR 0.5, 95% CI 0.2-1.0, and OR 1.0, 95% CI 0.5-2.1 for African-American and white women, respectively). For maternal age there was an approximately threefold increase in risk in women whose mothers were older than 22 years of age, relative to 19-22 years of age, when the women were born. After adjustment for maternal age, older paternal age increased risk in the oldest and youngest age categories (relative to 23-27 years of age at the woman's birth: OR 1.6, 95% CI 0.8-3.1 for age 15-22 years; OR 1.2, 95% CI 0.7-2.2 for age 28-34 years; and OR 1.5, 95% CI 0.7-3.2 for age 35-56 years). There was no association with older paternal age for white women alone. After adjustment for maternal age (265 cases, 224 controls), a birth order of fifth or higher relative to first had an inverse association with breast cancer for women younger than 49 years old (OR 0.6, 95% CI 0.3-1.3). **CONCLUSION:** Although the CIs are wide, these results lend support to

the possibility that the prenatal period is important for subsequent breast cancer risk, but they do not support the estrogen hypothesis as a unifying theory for the influence of this period.

PMID: 15535848 [PubMed - in process]

Display Abstract ▼ Show: 20 ▼ Sort ▼ Send to Text ▼

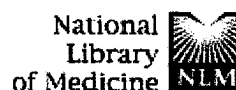
[Write to the Help Desk](#)

[NCBI](#) | [NLM](#) | [NIH](#)

[Department of Health & Human Services](#)

[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)

Mar 6 2005 07:57:22



My NCBI

[\[Sign In\]](#) [\[Register\]](#)

All Databases

PubMed

Nucleotide

Protein

Genome

Structure

OMIM

PMC

Journals

Books

Search PubMed for

Go Clear

☒ Limits Preview/Index History Clipboard Details

About Entrez

Field: Title/Abstract, Limits: Publication Date from 2000, English

Text Version

Display Abstract Show: 20 Sort Send to Text

All: 1 Review: 0

Entrez PubMed

Overview

Help | FAQ

Tutorial

New/Noteworthy

E-Utilities

PubMed Services

Journals Database

MeSH Database

Single Citation Matcher

Batch Citation Matcher

Clinical Queries

LinkOut

My NCBI (Cubby)

Related Resources

Order Documents

NLM Catalog

NLM Gateway

TOXNET

Consumer Health

Clinical Alerts

ClinicalTrials.gov

PubMed Central

1: J Urol. 2005 Apr;173(4):1246-51.

[Related Articles, Links](#)

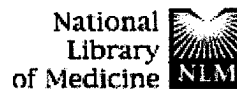
Structural assessment of the urethral sphincter in women with urinary retention.

Andrich DE, Rickards D, Landon DN, Fowler CJ, Mundy AR.

From the Institute of Urology, the Institute of Neurology (DNL, CJF), and The National Hospital for Neurology and Neurosurgery (CJF), Queen Square, London, United Kingdom.

PURPOSE:: The pathophysiology of urinary retention in women is generally unknown but a subgroup of women with urinary retention have been diagnosed as having so-called primary disorder of sphincter relaxation on the basis of an abnormal urethral sphincter electromyogram. It was suggested this sphincter overactivity could lead to work hypertrophy of the urethral rhabdosphincter and in this study we looked for any evidence of such muscle fiber hypertrophy.

MATERIALS AND METHODS:: In 9 women 18 to 45 years old (mean age 31.6) with urinary retention and overactive urethral sphincter electromyogram, light and electron microscopy were used to examine core needle biopsies of the urethral rhabdosphincter taken under transvaginal ultrasound control. Of the 9 patients only 5 biopsies processed for light microscopy and 4 processed for electron microscopy contained striated urethral muscle fibers. The results of these biopsies were compared to the morphology of a control specimen from a postmenopausal woman without a history of urinary retention. **RESULTS::** On light microscopy the urethral rhabdosphincter fiber diameter did not differ among patients (mean average 7.6 μ m), was less than that reported in the literature (15 to 20), but did not differ from that of the control (mean 9.9). In all patients electron microscopy showed excessive peripheral sarcoplasm with lipid and glycogen deposition, and sarcoplasmic accumulation of normal mitochondria. These ultrastructural abnormalities were not seen in the control. **CONCLUSIONS::** To our knowledge this is the first morphological description of the urethral rhabdosphincter in a subgroup of women with urinary retention. Mean rhabdosphincter fiber diameter was approximately the same in patients and controls. This study does not support the previous theory that urethral sphincter overactivity in a subgroup of women with urinary retention leads to work hyperplasia of urethral rhabdosphincter fibers. An alternative hypothesis is suggested.



My NCBI

[\[Sign In\]](#) [\[Register\]](#)

All Databases PubMed Nucleotide Protein Genome Structure OMIM PMC Journals Books

Search PubMed for Go Clear

☒ Limits Preview/Index History Clipboard Details

About Entrez

Field: Title/Abstract, Limits: Publication Date from 2000, English

Text Version

Display Abstract Show: 20 Sort Send to Text

All: 1 Review: 0

Entrez PubMed

Overview

Help | FAQ

Tutorial

New/Noteworthy

E-Utilities

PubMed Services

Journals Database

MeSH Database

Single Citation Matcher

Batch Citation Matcher

Clinical Queries

LinkOut

My NCBI (Cubby)

Related Resources

Order Documents

NLM Catalog

NLM Gateway

TOXNET

Consumer Health

Clinical Alerts

ClinicalTrials.gov

PubMed Central

1: J Theor Biol. 2005 May 21;234(2):277-88. Epub 2005 Jan 24. [Related Articles](#), [Links](#)

RESEARCH ARTICLE
FULL-TEXT ARTICLE

LEHIGH
S-F-X LINKS

On the importance of fatty acid composition of membranes for aging.

Hulbert AJ.

Metabolic Research Centre, University of Wollongong, Wollongong, NSW 2522, Australia; School of Biological Sciences, University of Wollongong, Wollongong, NSW 2522, Australia.

The membrane pacemaker theory of aging is an extension of the oxidative stress theory of aging. It emphasises variation in the fatty acid composition of membranes as an important influence on lipid peroxidation and consequently on the rate of aging and determination of lifespan. The products of lipid peroxidation are reactive molecules and thus potent damagers of other cellular molecules. It is suggested that the feedback effects of these peroxidation products on the oxidative stress experienced by cells is an important part of the aging process. The large variation in the chemical susceptibility of individual fatty acids to peroxidation coupled with the known differences in membrane composition between species can explain the different lifespans of species, especially the difference between mammals and birds as well as the body-size-related variation in lifespan within mammals and birds. Lifespan extension by calorie-restriction can also be explained by changes in membrane fatty acid composition which result in membranes more resistant to peroxidation. It is suggested that lifespan extension by reduced insulin/IGF signalling may also be mediated by changes in membrane fatty acid composition.

PMID: 15757684 [PubMed - in process]

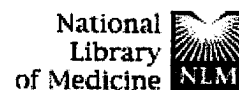
Display Abstract Show: 20 Sort Send to Text

[Write to the Help Desk](#)

[NCBI](#) | [NLM](#) | [NIH](#)

[Department of Health & Human Services](#)

[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)



My NCBI

[\[Sign In\]](#) [\[Register\]](#)[All Databases](#)[PubMed](#)[Nucleotide](#)[Protein](#)[Genome](#)[Structure](#)[OMIM](#)[PMC](#)[Journals](#)[Books](#)

Search

PubMed

for

Go

Clear

☒ Limits[Preview/Index](#)[History](#)[Clipboard](#)[Details](#)[About Entrez](#)[Text Version](#)[Entrez PubMed](#)[Overview](#)[Help | FAQ](#)[Tutorial](#)[New/Noteworthy](#)[E-Utilities](#)[PubMed Services](#)[Journals Database](#)[MeSH Database](#)[Single Citation Matcher](#)[Batch Citation Matcher](#)[Clinical Queries](#)[LinkOut](#)[My NCBI \(Cubby\)](#)[Related Resources](#)[Order Documents](#)[NLM Catalog](#)[NLM Gateway](#)[TOXNET](#)[Consumer Health](#)[Clinical Alerts](#)[ClinicalTrials.gov](#)[PubMed Central](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000**

Display

Abstract

Show: 20

Sort

Send to

Text

All: 1

Review: 0

1: J Antimicrob Chemother. 2005 Mar 10; [Epub ahead of print] [Related Articles](#), [Links](#)Full text article at
jac.oupjournals.org

Antitubercular inhaled therapy: opportunities, progress and challenges.

Pandey R, Khuller GK.

Department of Biochemistry, Postgraduate Institute of Medical Education & Research, Chandigarh--160 012, India.

Pulmonary tuberculosis remains the commonest form of this disease and the development of methods for delivering antitubercular drugs directly to the lungs via the respiratory route is a rational therapeutic goal. The obvious advantages of inhaled therapy include direct drug delivery to the diseased organ, targeting to alveolar macrophages harbouring the mycobacteria, reduced risk of systemic toxicity and improved patient compliance. Research efforts have demonstrated the feasibility of various drug delivery systems employing liposomes, polymeric microparticles and nanoparticles to serve as inhalable antitubercular drug carriers. In particular, nanoparticles have emerged as a remarkably useful tool for this purpose. While some researchers have preferred dry powder inhalers, others have emphasized nebulization. Beginning with the respiratory delivery of a single antitubercular drug, it is now possible to deliver multiple drugs simultaneously with a greater therapeutic efficacy. More experience and expertise have been observed with synthetic polymers, nevertheless, the possibility of using natural polymers for inhaled therapy has yet to be explored. Several key issues such as patient education, cost of treatment, stability and large scale production of drug formulations, etc. need to be addressed before antitubercular inhaled therapy finds its way from theory to clinical reality.

PMID: 15761077 [PubMed - as supplied by publisher]

Display

Abstract

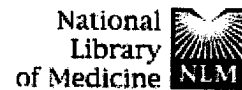
Show: 20

Sort

Send to

Text

[Write to the Help Desk](#)[NCBI | NLM | NIH](#)[Department of Health & Human Services](#)



My NCBI

[\[Sign In\]](#) [\[Register\]](#)

All Databases

PubMed

Nucleotide

Protein

Genome

Structure

OMIM

PMC

Journals

Books

Search

PubMed

for

Go

Clear

☒ Limits

Preview/Index

History

Clipboard

Details

About Entrez

Text Version

Entrez PubMed

Overview

Help | FAQ

Tutorial

New/Noteworthy

E-Utilities

PubMed Services

Journals Database

MeSH Database

Single Citation Matcher

Batch Citation Matcher

Clinical Queries

LinkOut

My NCBI (Cubby)

Related Resources

Order Documents

NLM Catalog

NLM Gateway

TOXNET

Consumer Health

Clinical Alerts

ClinicalTrials.gov

PubMed Central

Field: Title/Abstract, Limits: Publication Date from 2000, English

Display

Abstract

Show: 20

Sort

Send to

Text

All: 1

Review: 1



1: Philos Trans R Soc Lond B Biol Sci. 2001 Jul
29;356(1411):1057-68.

Related Articles

Links

MetaPress

Early effects of climate change: do they include changes in vector-borne disease?

Kovats RS, Campbell-Lendrum DH, McMichael AJ, Woodward A, Cox JS.

Department of Epidemiology and Population Health, London School of Hygiene and Tropical Medicine, Keppel Street, London WC1E 7HT, UK.

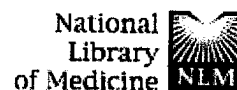
sari.kovats@lshtm.ac.uk

The world's climate appears now to be changing at an unprecedented rate. Shifts in the distribution and behaviour of insect and bird species indicate that biological systems are already responding to this change. It is well established that climate is an important determinant of the spatial and temporal distribution of vectors and pathogens. In theory, a change in climate would be expected to cause changes in the geographical range, seasonality (intra-annual variability), and in the incidence rate (with or without changes in geographical or seasonal patterns). The detection and then attribution of such changes to climate change is an emerging task for scientists. We discuss the evidence required to attribute changes in disease and vectors to the early effects of anthropogenic climate change. The literature to date indicates that there is a lack of strong evidence of the impact of climate change on vector-borne diseases (i.e. malaria, dengue, leishmaniasis, tick-borne diseases). New approaches to monitoring, such as frequent and long-term sampling along transects to monitor the full latitudinal and altitudinal range of specific vector species, are necessary in order to provide convincing direct evidence of climate change effects. There is a need to reassess the appropriate levels of evidence, including dealing with the uncertainties attached to detecting the health impacts of global change.

Publication Types:

- Review
- Review, Tutorial

PMID: 11516383 [PubMed - indexed for MEDLINE]



My NCBI

[\[Sign In\]](#) [\[Register\]](#)[All Databases](#)[PubMed](#)[Nucleotide](#)[Protein](#)[Genome](#)[Structure](#)[OMIM](#)[PMC](#)[Journals](#)[Books](#)

Search PubMed

▼ for

Go

Clear



Limits

[Preview/Index](#)[History](#)[Clipboard](#)[Details](#)[About Entrez](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000, English**

Text Version

Display

Abstract ▼

Show: 20 ▼

Sort ▼

Send to

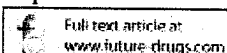
Text ▼

All: 1

Review: 0

[Entrez PubMed](#)[Overview](#)[Help | FAQ](#)[Tutorial](#)[New/Noteworthy](#)[E-Utilities](#)[PubMed Services](#)[Journals Database](#)[MeSH Database](#)[Single Citation Matcher](#)[Batch Citation Matcher](#)[Clinical Queries](#)[LinkOut](#)[My NCBI \(Cubby\)](#)[Related Resources](#)[Order Documents](#)[NLM Catalog](#)[NLM Gateway](#)[TOXNET](#)[Consumer Health](#)[Clinical Alerts](#)[ClinicalTrials.gov](#)[PubMed Central](#)

1: Expert Rev Vaccines. 2005 Feb;4(1):51-62.

[Related Articles, Links](#)

DNA vaccines for poultry: the jump from theory to practice.

Haygreen L, Davison F, Kaiser P.

Institute for Animal Health, Compton, Berkshire RG20 7NN, UK.

DNA vaccines could offer a solution to a number of problems faced by the poultry industry; they are relatively easy to manufacture, stable, potentially easy to administer, can overcome neonatal tolerance and the deleterious effects of maternal antibody, and do not cause disease pathology. Combined with this, in ovo vaccination offers the advantage of reduced labor costs, mass administration and the induction of an earlier immune response. Together, this list of advantages is impressive. However, this combined technology is still in its infancy and requires many improvements. The potential of CpG motifs, DNA vaccines and in ovo vaccination, however, can be observed by the increasing number of recent reports investigating their application in challenge experiments. CpG motifs have been demonstrated to be stimulatory both invitro and invivo. In addition, DNA vaccines have been successfully delivered via the in ovo route, albeit not yet through the amniotic fluid. Lastly, a recent report has demonstrated that a DNA vaccine against infectious bronchitis virus administered via in ovo vaccination, followed by live virus boost, can slightly improve on the protective effect induced by the live virus alone. Therefore, DNA vaccination via the in ovo route is promising and offers potential as a poultry vaccine, however, efficacy needs to be improved and the costs of production reduced before it is likely to be beneficial to the poultry industry in the long term.

PMID: 15757473 [PubMed - in process]

Display

Abstract ▼

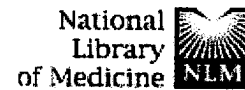
Show: 20 ▼

Sort ▼

Send to

Text ▼

[Write to the Help Desk](#)[NCBI | NLM | NIH](#)[Department of Health & Human Services](#)[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)



My NCBI

[\[Sign In\]](#) [\[Register\]](#)

All Databases

PubMed

Nucleotide

Protein

Genome

Structure

OMIM

PMC

Journals

Books

Search PubMed

▼ for

Go

Clear

☒ Limits[Preview/Index](#)[History](#)[Clipboard](#)[Details](#)[About Entrez](#)[Text Version](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000, English**[Display](#)[Abstract](#)

▼ Show: 20

[Sort](#)

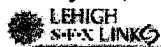
▼

[Send to](#)[Text](#)

▼

All: 1

Review: 0

☐ 1: J Theor Biol. 2005 May 21;234(2):167-72. Epub 2005 Jan 20. [Related Articles](#), [Links](#)

Daisyworld inhabited with daisies incorporating a seed size/number trade-off: the mechanism of negative feedback on selection from a standpoint of the competition theory.

Seto M, Akagi T.

Faculty of Agriculture, Tokyo University of Agriculture and Technology, 3-5-8 Saiwai-cho, Fuchu-shi, Tokyo 183-8509, Japan.

We reexamined a Daisyworld model from the traditional view of competition theory. Unlike the original model, white and black daisies in our model incorporate a seeding/germination trade-off against bare ground area without assuming the local temperature reward. As a result, the planetary temperature is automatically regulated by two species if the following conditions are met: (i) the species react equally to an environmental condition, but one can alter the environmental condition in the opposite direction to the other. (ii) that one of the two cannot have both a higher maximal growth rate ($\mu(\max)$) and lower half-saturation constant (K) than those of the other. In other words, a pair of phenotypes incorporates a trade-off between quality and number of seeds. We found that the homeostatic regulation can also be reconciled with the adaptive evolution of optimal temperature. The results of simulation imply that biotic environmental feedback can also be maintained when the emergence of polymorphisms (black and white daisies) is closely linked to such a trade-off.

PMID: 15757676 [PubMed - in process]

[Display](#)[Abstract](#)

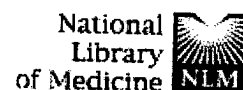
Show: 20

[Sort](#)[Send to](#)[Text](#)[Write to the Help Desk](#)[NCBI](#) | [NLM](#) | [NIH](#)

Department of Health & Human Services

[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)

Mar 6 2005 07:57:22



My NCBI

[\[Sign In\]](#) [\[Register\]](#)

All Databases

PubMed

Nucleotide

Protein

Genome

Structure

OMIM

PMC

Journals

Books

Search PubMed

for

Go

Clear

☒ Limits[Preview/Index](#)[History](#)[Clipboard](#)[Details](#)[About Entrez](#)[Text Version](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000, English**[Display](#)

Abstract

Show: 20

Sort

Send to

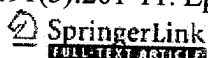
Text

All: 1

Review: 0

[Entrez PubMed](#)[Overview](#)[Help | FAQ](#)[Tutorial](#)[New/Noteworthy](#)[E-Utilities](#)[PubMed Services](#)[Journals Database](#)[MeSH Database](#)[Single Citation Matcher](#)[Batch Citation Matcher](#)[Clinical Queries](#)[LinkOut](#)[My NCBI \(Cubby\)](#)[Related Resources](#)[Order Documents](#)[NLM Catalog](#)[NLM Gateway](#)[TOXNET](#)[Consumer Health](#)[Clinical Alerts](#)[ClinicalTrials.gov](#)[PubMed Central](#)

1: J Comp Physiol A Neuroethol Sens Neural Behav Physiol. 2005 Mar;191(3):201-11. Epub 2004 Oct 12.

[Related Articles,
Links](#)

Expectations and outcomes: decision-making in the primate brain.

McCoy AN, Platt ML.

Department of Neurobiology, Duke University Medical Center, 325 Bryan Research Building, Box 3209, Durham, NC, 27710, USA.

Success in a constantly changing environment requires that decision-making strategies be updated as reward contingencies change. How this is accomplished by the nervous system has, until recently, remained a profound mystery. New studies coupling economic theory with neurophysiological techniques have revealed the explicit representation of behavioral value. Specifically, when fluid reinforcement is paired with visually-guided eye movements, neurons in parietal cortex, prefrontal cortex, the basal ganglia, and superior colliculus-all nodes in a network linking visual stimulation with the generation of oculomotor behavior-encode the expected value of targets lying within their response fields. Other brain areas have been implicated in the processing of reward-related information in the abstract: midbrain dopaminergic neurons, for instance, signal an error in reward prediction. Still other brain areas link information about reward to the selection and performance of specific actions in order for behavior to adapt to changing environmental exigencies. Neurons in posterior cingulate cortex have been shown to carry signals related to both reward outcomes and oculomotor behavior, suggesting that they participate in updating estimates of orienting value.

PMID: 15759141 [PubMed - in process]

Display

Abstract

Show: 20

Sort

Send to

Text

[Write to the Help Desk](#)[NCBI | NLM | NIH](#)[Department of Health & Human Services](#)[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)

[About Entrez](#)[Text Version](#)[Entrez PubMed](#)[Overview](#)[Help | FAQ](#)[Tutorial](#)[New/Noteworthy](#)[E-Utilities](#)[PubMed Services](#)[Journals Database](#)[MeSH Database](#)[Single Citation Matcher](#)[Batch Citation Matcher](#)[Clinical Queries](#)[LinkOut](#)[My NCBI \(Cubby\)](#)[Related Resources](#)[Order Documents](#)[NLM Catalog](#)[NLM Gateway](#)[TOXNET](#)[Consumer Health](#)[Clinical Alerts](#)[ClinicalTrials.gov](#)[PubMed Central](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000**[Display](#)[Abstract](#)

Show: 20

[Sort](#)[Send to](#)[Text](#)

All: 1

Review: 0



1: J Neurol Sci. 2005 Mar 15;229-230(1):109-16. Epub 2004 Dec

[Related Articles](#)

8.

[Links](#)

Effects of arousing emotional scenes on the distribution of visuospatial attention: changes with aging and early subcortical vascular dementia.

Rosler A, Ulrich C, Billino J, Sterzer P, Weidauer S, Bernhardt T, Steinmetz H, Frolich L, Kleinschmidt A.

Department of Neurology, Johann Wolfgang Goethe-University, Frankfurt, Germany.

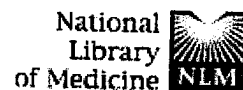
BACKGROUND: The modulation of attention by emotionally arousing stimuli is highly important for each individual's social function. Disturbances of emotional processing are a supportive feature for the diagnosis of subcortical vascular dementia (SVD). We address here whether these disturbances might be useful as an early disease marker. **METHODS:** In order to examine the modulation of visual attention by emotionally arousing stimuli of different valence, 12 elderly patients with early SVD, 12 age-comparable healthy adults and 12 young healthy subjects were studied while looking at pairs of pictures from the International Affective Picture Battery that were either neutral-neutral, neutral-positive or neutral-negative in terms of emotional content. Eye movements were recorded with an infrared eye-tracking system. The direction of the first saccade and the dwell time during the 10 s of presentation were measured and compared among groups with parametric tests. **RESULTS:** All subjects showed a modulation of initial attentional orienting as well as a higher percentage of dwell time towards the pictures containing emotional material. Patients with SVD and old controls did not differ in either experimental measure. Young patients showed a stronger bias towards emotionally negative material than both groups of older individuals. **CONCLUSIONS:** Modulation of visuospatial attention is preserved in early SVD. This might have implications for therapeutic interventional approaches. A weakened sustained attention towards negative but not positive emotional pictures in the elderly is in accordance with the socioemotional selectivity theory describing a relative selection of positive stimuli with aging.

PMID: 15760628 [PubMed - in process]

[Display](#)[Abstract](#)

Show: 20

[Sort](#)[Send to](#)[Text](#)[Write to the Help Desk](#)[NCBI](#) | [NLM](#) | [NIH](#)[Department of Health & Human Services](#)[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)



My NCBI

[\[Sign In\]](#) [\[Register\]](#)

All Databases

PubMed

Nucleotide

Protein

Genome

Structure

OMIM

PMC

Journals

Books

Search PubMed

for

Go

Clear

☒ Limits[Preview/Index](#)[History](#)[Clipboard](#)[Details](#)[About Entrez](#)Field: **Title/Abstract**, Limits: **Publication Date from 2000**

Text Version

[Display](#)[Abstract](#)

Show: 20

Sort

Send to [Text](#)

All: 1

Review: 0

[Entrez PubMed](#)[Overview](#)[Help | FAQ](#)[Tutorial](#)[New/Noteworthy](#)[E-Utilities](#)[PubMed Services](#)[Journals Database](#)[MeSH Database](#)[Single Citation Matcher](#)[Batch Citation Matcher](#)[Clinical Queries](#)[LinkOut](#)[My NCBI \(Cubby\)](#)[Related Resources](#)[Order Documents](#)[NLM Catalog](#)[NLM Gateway](#)[TOXNET](#)[Consumer Health](#)[Clinical Alerts](#)[ClinicalTrials.gov](#)[PubMed Central](#)☐ 1: Hum Reprod. 2005 Mar 10; [Epub ahead of print][Related Articles, Links](#)Full text article at
humrep.oupjournals.org

New debate: natural killer cells and reproductive failure--theory, practice and prejudice.

Rai R, Sacks G, Trew G.

Department of Obstetrics and Gynaecology, Faculty of Medicine, Imperial College London, St Mary's Hospital, London, UK.

The relationship between peripheral blood natural killer (NK) cells and reproductive failure is one of the most controversial areas in reproductive medicine. Amidst much publicity, peripheral blood NK cell testing is being promoted as a useful diagnostic test to guide the initiation of a variety of immunosuppressive therapies amongst patients with either recurrent miscarriage or infertility. We contend (i) that at present there is no scientific basis for the introduction of NK cell testing into routine clinical practice, and (ii) that the use of immunosuppressant agents based on the results of such testing may potentially be harmful.

PMID: 15760961 [PubMed - as supplied by publisher]

[Display](#)[Abstract](#)

Show: 20

Sort

Send to [Text](#)[Write to the Help Desk](#)[NCBI](#) | [NLM](#) | [NIH](#)[Department of Health & Human Services](#)[Privacy Statement](#) | [Freedom of Information Act](#) | [Disclaimer](#)

Mar 6 2005 07:57:22

Exhibit 2

A drawing of the bacterial
flagellum from a biochemistry
textbook

The bacterial flagellum

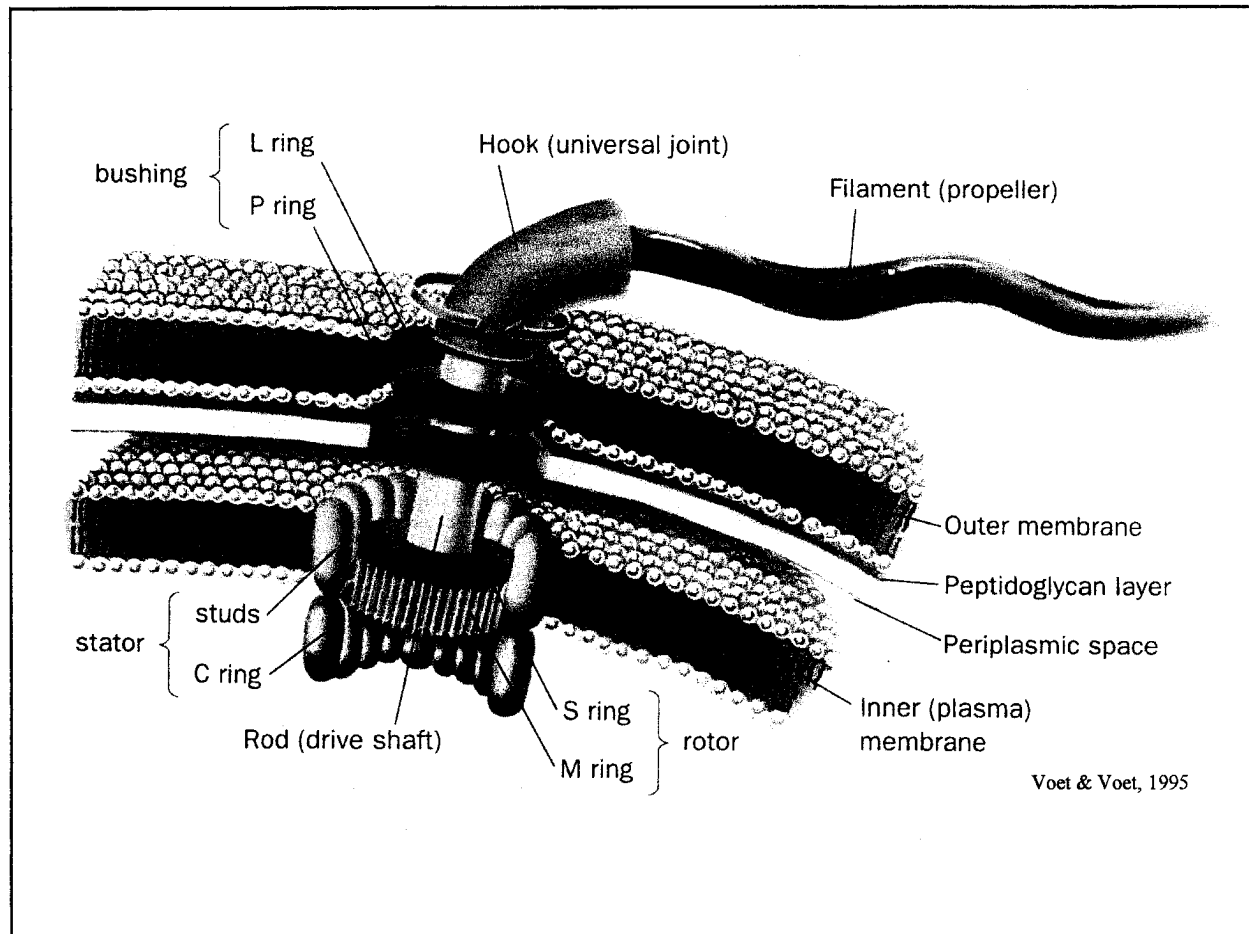
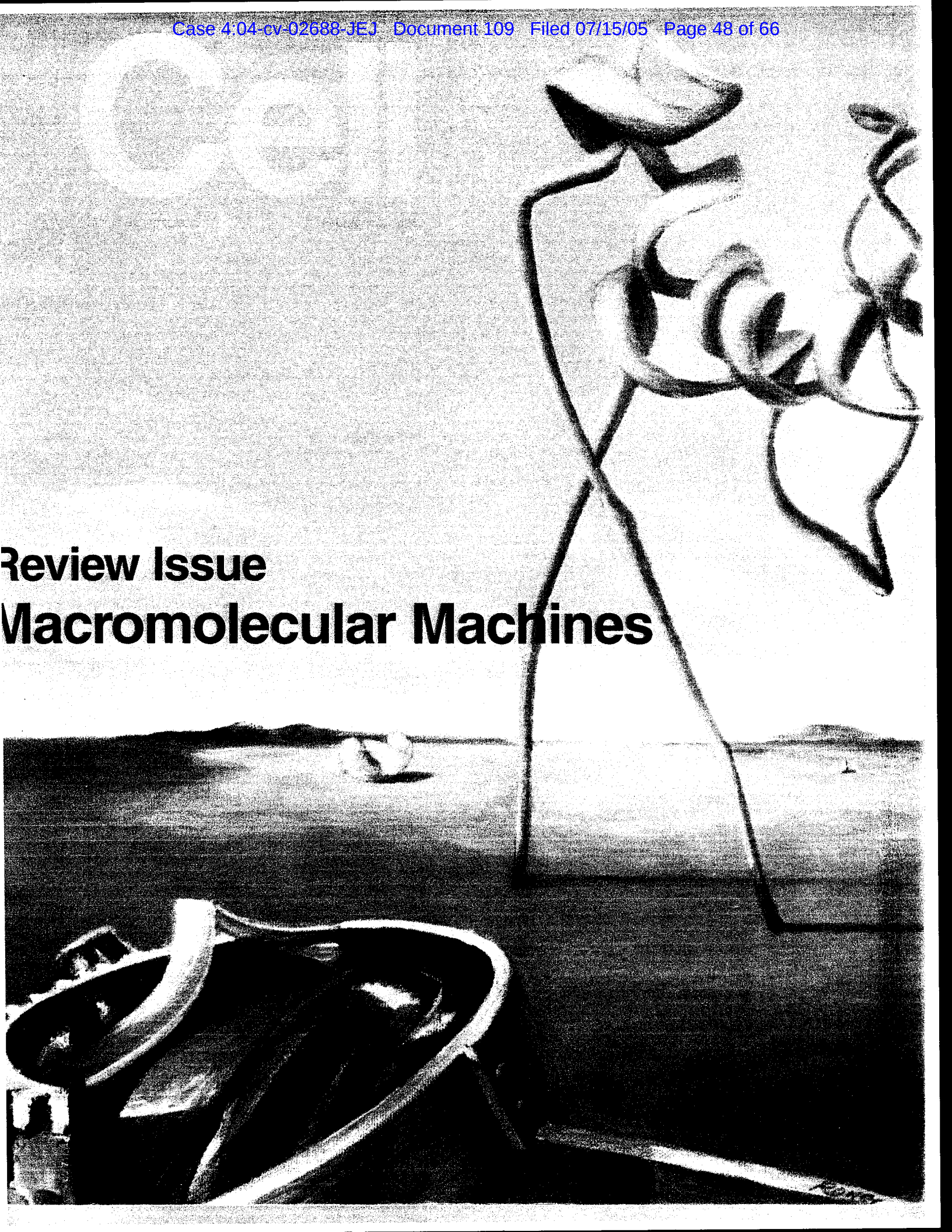


Exhibit 3

Cover and Table of Contents of
the February 6, 1998 issue of
the journal *Cell*



Review Issue

Macromolecular Machines

Volume 92 Number 3 February 6, 1998

Cell

Reviews on Macromolecular Machines

The Cell as a Collection of Protein Machines: Preparing the Next Generation of Molecular Biologists	B. Alberts	291
Polymerases and the Replisome: Machines within Machines	T. A. Baker and S. P. Bell	295
Eukaryotic Transcription: An Interlaced Network of Transcription Factors and Chromatin-Modifying Machines	J. T. Kadonaga	307
Mechanical Devices of the Spliceosome: Motors, Clocks, Springs, and Things	J. P. Staley and C. Guthrie	315
Nucleocytoplasmic Transport: The Last 200 Nanometers	M. Ohno, M. Fornerod, and I. W. Mattaj	327
Molecular Movement inside the Translational Engine	K. S. Wilson and H. F. Noller	337
The Hsp70 and Hsp60 Chaperone Machines	B. Bukau and A. L. Horwich	351
The Proteasome: Paradigm of a Self-Compartmentalizing Protease	W. Baumeister, J. Walz, F. Zühl, and E. Seemüller	367
Protein Translocation: Tunnel Vision	K. E. S. Matlack, W. Mothes, and T. A. Rapoport	381

(continued)

Articles

Disruption of Angiogenesis by PEX, a Noncatalytic Metalloproteinase Fragment with Integrin Binding Activity	P. C. Brooks, S. Silletti, T. L. von Schalscha, M. Friedlander, and D. A. Cheresh	391
TRF2 Protects Human Telomeres from End-to-End Fusions	B. van Steensel, A. Smogorzewska, and T. de Lange	401
Structure of the Guanine Nucleotide Exchange Factor Sec7 Domain of Human Arno and Analysis of the Interaction with ARF GTPase	E. Mossessova, J. M. Gulbis, and J. Goldberg	415

Positions Available

Conferences and Courses

Like the machines invented by humans to deal efficiently with the macroscopic world, protein assemblies contain highly coordinated moving parts. Reviewed in this issue of *Cell* are the protein machines that control replication, transcription, splicing, nucleocytoplasmic transport, protein synthesis, protein assembly, protein degradation, and protein translocation—the machines that underlie the workings of all living things. The cover, inspired by the idea of protein machines, is a reproduction of the painting “Dilemma of the Helixes” by Rong Li, Ph.D., Department of Cell Biology, Harvard Medical School.

Exhibit 4

New York Times op-ed
summarizing the argument for
intelligent design

LexisNexis

[Home](#) [Sources](#) [How Do I?](#) [Site Map](#) [What's New](#) [Help](#)

Search Terms: behe

FOCUS™

[Search Within Results](#)[Edit Search](#)[Print](#) [Email](#)

Document 1 of 1.

Full

Copyright 2005 The **New York Times** Company
The **New York Times**

February 7, 2005 Monday
Late Edition - Final

SECTION: Section A; Column 1; Editorial Desk; Pg. 21**LENGTH:** 965 words**HEADLINE:** Design for Living**BYLINE:** By Michael J. **Behe**.

Michael J. **Behe**, a professor of biological sciences at Lehigh University and a senior fellow with the Discovery Institute's Center for Science and Culture, is the author of "Darwin's Black Box: The Biochemical Challenge to Evolution."

DATELINE: BETHLEHEM, Pa.**BODY:**

IN the wake of the recent lawsuits over the teaching of Darwinian evolution, there has been a rush to debate the merits of the rival theory of intelligent design. As one of the scientists who have proposed design as an explanation for biological systems, I have found widespread confusion about what intelligent design is and what it is not.

First, what it isn't: the theory of intelligent design is not a religiously based idea, even though devout people opposed to the teaching of evolution cite it in their arguments. For example, a critic recently caricatured intelligent design as the belief that if evolution occurred at all it could never be explained by Darwinian natural selection and could only have been directed at every stage by an omniscient creator. That's misleading. Intelligent design proponents do question whether random mutation and natural selection completely explain the deep structure of life. But they do not doubt that evolution occurred. And intelligent design itself says nothing about the religious concept of a creator.

Rather, the contemporary argument for intelligent design is based on physical evidence and a straightforward application of logic. The argument for it consists of four linked claims. The first claim is uncontroversial: we can often recognize the effects of design in nature. For example, unintelligent physical forces like plate tectonics and erosion seem quite sufficient to account for the origin of the Rocky Mountains. Yet they are not enough to explain Mount Rushmore.

Of course, we know who is responsible for Mount Rushmore, but even someone who had never heard of the monument could recognize it as designed. Which leads to the second claim of the intelligent design argument: the physical marks of design are visible in aspects of biology. This is uncontroversial, too. The 18th-century clergyman William Paley likened living things to a watch, arguing that the workings of both point to intelligent design. Modern Darwinists disagree with Paley that the perceived design is real, but they do agree that life overwhelms us with the appearance of design.

For example, Francis Crick, co-discoverer of the structure of DNA, once wrote that biologists must constantly remind themselves that what they see was not designed but evolved. (Imagine a scientist repeating through clenched teeth: "It wasn't really designed. Not really.")

The resemblance of parts of life to engineered mechanisms like a watch is enormously stronger than what Reverend Paley imagined. In the past 50 years modern science has shown that the cell, the very foundation of life, is run by machines made of molecules. There are little molecular trucks in the cell to ferry supplies, little outboard motors to push a cell through liquid.

In 1998 an issue of the journal *Cell* was devoted to molecular machines, with articles like "The Cell as a Collection of Protein Machines" and "Mechanical Devices of the Spliceosome: Motors, Clocks, Springs and Things." Referring to his student days in the 1960's, Bruce Alberts, president of the National Academy of Sciences, wrote that "the chemistry that makes life possible is much more elaborate and sophisticated than anything we students had ever considered." In fact, Dr. Alberts remarked, the entire cell can be viewed as a factory with an elaborate network of interlocking assembly lines, each of which is composed of a set of large protein machines. He emphasized that the term machine was not some fuzzy analogy; it was meant literally.

The next claim in the argument for design is that we have no good explanation for the foundation of life that doesn't involve intelligence. Here is where thoughtful people part company. Darwinists assert that their theory can explain the appearance of design in life as the result of random mutation and natural selection acting over immense stretches of time. Some scientists, however, think the Darwinists' confidence is unjustified. They note that although natural selection can explain some aspects of biology, there are no research studies indicating that Darwinian processes can make molecular machines of the complexity we find in the cell.

Scientists skeptical of Darwinian claims include many who have no truck with ideas of intelligent design, like those who advocate an idea called complexity theory, which envisions life self-organizing in roughly the same way that a hurricane does, and ones who think organisms in some sense can design themselves.

The fourth claim in the design argument is also controversial: in the absence of any convincing non-design explanation, we are justified in thinking that real intelligent design was involved in life. To evaluate this claim, it's important to keep in mind that it is the profound appearance of design in life that everyone is laboring to explain, not the appearance of natural selection or the appearance of self-organization.

The strong appearance of design allows a disarmingly simple argument: if it looks, walks and quacks like a duck, then, absent compelling evidence to the contrary, we have warrant to conclude it's a duck. Design should not be overlooked simply because it's so obvious.

Still, some critics claim that science by definition can't accept design, while others argue that science should keep looking for another explanation in case one is out there. But we can't settle questions about reality with definitions, nor does it seem useful to search relentlessly for a non-design explanation of Mount Rushmore. Besides, whatever special restrictions scientists adopt for themselves don't bind the public, which polls show, overwhelmingly, and sensibly, thinks that life was designed. And so do many scientists who see roles for both the messiness of evolution and the elegance of design.

URL: <http://www.nytimes.com>

GRAPHIC: Drawing (Drawing by Leigh Wells)

LOAD-DATE: February 7, 2005

Document 1 of 1.

[Terms & Conditions](#) [Privacy](#) [Copyright](#) © 2005 LexisNexis, a division of Reed Elsevier Inc. All Rights Reserved.

Exhibit 5

**“Directed Panspermia”, by
Francis Crick and Leslie Orgel**

ICARUS 19, 341-346 (1973)

Directed Panspermia

F. H. C. CRICK

Medical Research Council, Laboratory of Molecular Biology, Hills Road, Cambridge, England

AND

L. E. ORGEL

The Salk Institute for Biological Studies, P.O. Box 1809, San Diego, California 92112

Received June 22, 1972; revised December 20, 1972

[It now seems unlikely that extraterrestrial living organisms could have reached the earth either as spores driven by the radiation pressure from another star or as living organisms imbedded in a meteorite. As an alternative to these nineteenth-century mechanisms, we have considered Directed Panspermia, the theory that organisms were deliberately transmitted to the earth by intelligent beings on another planet.] We conclude that it is possible that life reached the earth in this way, but that the scientific evidence is inadequate at the present time to say anything about the probability. We draw attention to the kinds of evidence that might throw additional light on the topic.

INTRODUCTION

It was not until the middle of the nineteenth century that Pasteur and Tyndall completed the demonstration that spontaneous generation is not occurring on the Earth nowadays. Darwin and a number of other biologists concluded that life must have evolved here long ago when conditions were more favourable. A number of scientists, however, drew a quite different conclusion. They supposed that if life does not evolve from terrestrial nonliving matter nowadays, it may never have done so. Hence, they argued, life reached the earth as an "infection" from another planet (Oparin, 1957).

Arrhenius (1908) proposed that spores had been driven here by the pressure of the light from the central star of another planetary system. His theory is known as Panspermia. Kelvin suggested that the first organisms reached the Earth in a meteorite. Neither of these theories is absurd, but both can be subjected to severe criticism. Sagan (Shklovski and Sagan, 1966; Sagan and Whitehall, 1973) has shown that any known type of radiation-

resistant spore would receive so large a dose of radiation during its journey to the Earth from another Solar System that it would be extremely unlikely to remain viable. The probability that sufficiently massive objects escape from a Solar System and arrive on the planet of another one is considered to be so small that it is unlikely that a single meteorite of extrasolar origin has ever reached the surface of the Earth (Sagan, private communication). These arguments may not be conclusive, but they argue against the "infective" theories of the origins of life that were proposed in the nineteenth century.

It has also been argued that "infective" theories of the origins of terrestrial life should be rejected because they do no more than transfer the problem of origins to another planet. This view is mistaken; the historical facts are important in their own right. For all we know there may be other types of planet on which the origin of life *ab initio* is greatly more probable than on our own. For example, such a planet may possess a mineral, or compound, of crucial catalytic importance, which is rare on

liberately
ologically
ret.
d libe
ness: It
weak and
stars, of
an, have
f which
rm of life
d liquid
tudies of
als under
m likely
ates on a
planets,
nothing
evolves
a soup,
still less
ves in a
may be
be alone
igh the
life of
re thus
en (and
where
least a
n years
ved to
atural
exist,
ground
e that
would
would
forms
fe de-
of an
been
ore or
scien-
16
olved
ently

left behind by some previous visitors from another planet (for example, in their garbage). Here we wish to examine a very specific form of Directed Panspermia. Could life have started on Earth as a result of infection by microorganisms sent here deliberately by a technological society on another planet, by means of a special long-range unmanned spaceship? To show that this is not totally implausible we shall use the theorem of detailed cosmic reversibility; if we are capable of infecting an as yet lifeless extrasolar planet, then, given that the time was available, another technological society might well have infected our planet when it was still lifeless.

THE PROPOSED SPACESHIP

The spaceship would carry large samples of a number of microorganisms, each having different but simple nutritional requirements, for example blue-green algae, which could grow on CO_2 and water in "sunlight." A payload of 1000 kg might be made up of 10 samples each containing 10^{16} microorganisms, or 100 samples each of 10^{15} microorganisms.

It would not be necessary to accelerate the spaceship to extremely high velocities, since its time of arrival would not be important. The radius of our galaxy is about 10^5 light years, so we could infect most planets in the galaxy within 10^8 yr by means of a spaceship traveling at only one-thousandth of the velocity of light. Several thousand stars are within a hundred light years of the Earth and could be reached within as little as a million years by a spaceship travelling at only 60,000 mph, or within 10,000 yr if a speed of one-hundredth of that of light were possible.

The technology required to carry out such an act of interstellar pollution is not available at the present time. However, it seems likely that the improvements in astronomical techniques will permit the location of extrasolar planets within the next few decades. Similarly, the problem of sending spaceships to other stars, at velocities low compared with that of light, should not prove insoluble once workable

nuclear engines are available. This again is likely to be within a few decades. The most difficult problem would be presented by the long flight times; it is not clear how long it will be before we can build components that would survive in space for periods of thousands or millions of years.

Although there are some technological problems associated with the distribution of the microorganisms in viable form after a long journey through space, none of them seems insuperable. Some radiation protection could be provided during the journey. Suitable packaging should guarantee that small samples, including some viable organisms, would be widely distributed. The question of how long microorganisms, and in particular bacterial spores, could survive in a spaceship has been considered in a preliminary way by Sneath (1962). He concludes "that life could probably be preserved for periods of more than a million years if suitably protected and maintained at temperatures close to absolute zero." Sagan (1960) has given a comparable estimate of the effects of radiation damage. We conclude that within the foreseeable future we could, if we wished, infect another planet, and hence that it is not out of the question that our planet was infected.

We can in fact go further than this. It may be possible in the future to send either mice or men or elaborate instruments to the planets of other Solar Systems (as so often described in science fiction) but a rocket carrying microorganisms will always have a much greater effective range and so be advantageous if the sole aim is to spread life. This is true for several reasons. The conditions on many planets are likely to favour microorganisms rather than higher organisms. Because of their extremely small size vast numbers of microorganisms can be carried, so much more wastage can be accepted. The ability of microorganisms to survive, without special equipment, both storage for very long periods at low temperatures and also an abrupt change back to room temperatures is also a great advantage. Whatever the potential range for infection by other organisms, microorganisms can almost

began, only a single one might have operated in the organisms used to infect the Earth.

CONCLUSION

In summary, there is adequate time for technological society to have evolved twice in succession. The places in the galaxy where life could start, if seeded, are probably very numerous. We can foresee that we ourselves will be able to construct rockets with sufficient range, delivery ability, and surviving payload if microorganisms are used. Thus the idea of Directed Panspermia cannot at the moment be rejected by any simple argument. It is radically different from the idea that life started here *ab initio* without infection from elsewhere. We have thus two sharply different theories of the origin of life on Earth. Can we choose between them?

At the moment it seems that the experimental evidence is too feeble to make this discrimination. It is difficult to avoid a personal prejudice, one way or the other, but such prejudices find no scientific support of any weight. It is thus important that both theories should be followed up. Work on the supposed terrestrial origin of life is in progress in many laboratories. As far as Directed Panspermia is concerned we can suggest several rather diverse lines of research.

The arguments we have employed here are, of necessity, somewhat sketchy. Thus the detailed design of a long-range spaceship would be worth a careful feasibility study. The spaceship must clearly be able to home on a star, for an object with any appreciable velocity, if dispatched in a random direction, would in almost all cases pass right through the galaxy and out the other side. It must probably have to decelerate as it approached the star, in order to allow the safe delivery of the payload. The packets of microorganisms must be made and dispersed in such a way that they can survive the entry at high velocity into the atmosphere of the planet, and yet be able to dissolve in the oceans. Many useful feasibility studies could be carried out on the engineering points involved.

On the biological side we lack precise

information concerning the life-time of microorganisms held at very low temperatures while traveling through space at relatively high velocities. The rocket would presumably be coasting most of the time so the convenient temperature might approximate to that of space. How serious is radiation damage, given a certain degree of shielding? How many distinct types of organism should be sent and which should they be? Should they collectively be capable of nitrogen fixation, oxidative phosphorylation and photosynthesis? Although many "soups" have been produced artificially in the laboratory, following the pioneer experiments of Miller, as far as we know no careful study has been made to determine which present-day organisms would grow well in them under primitive Earth conditions.

At the same time present-day organisms should be carefully scrutinized to see if they still bear any vestigial traces of extra-terrestrial origin. We have already mentioned the uniformity of the genetic code and the anomalous abundance of molybdenum. These facts amount to very little by themselves but as already stated there may be other as yet unsuspected features which, taken together, might point to a special type of planet as the home of our ancestors.

These enquiries are not trivial, for if successful they could lead to others which would touch us more closely. Are the senders or their descendants still alive? Or have the hazards of 4 billion years been too much for them? Has their star inexorably warmed up and frizzled them, or were they able to colonise a different Solar System with a short-range spaceship? Have they perhaps destroyed themselves, either by too much aggression or too little? The difficulties of placing any form of life on another planetary system are so great that we are unlikely to be their sole descendants. Presumably they would have made many attempts to infect the galaxy. If the range of their rockets were small this might suggest that we have cousins on planets which are not too distant. Perhaps the galaxy is lifeless except for a local village, of which we are one member.

Exhibit 6

“Biology Text illustrations More Fiction Than Fact”, New York Times, 4/8/01 (sidebar)

The New York Times

Copyright © 2001 The New York Times

SUNDAY, APRIL 8, 2001

VOL. CL . . . No. 51,717

Darwin vs. Design: Evolutionists' New Battle

By JAMES GLANZ

When Kansas school officials restored the theory of evolution to statewide education standards a few weeks ago, biologists might have been inclined to declare victory over creationism.

Instead, some evolutionists say, the latter stages of the battle in Kansas, along with new efforts in Michigan and Pennsylvania as well as in a number of universities and even in Washington, suggest that the issue is far from settled.

This time, though, the evolutionists find themselves arrayed not against traditional creationism, with its roots in biblical literalism, but against a more sophisticated idea: the intelligent design theory.

Proponents of this theory, led by a group of academics and intellectuals and including some biblical creationists, accept that the earth is billions of years old, not the thousands of years suggested by a literal reading of the Bible.

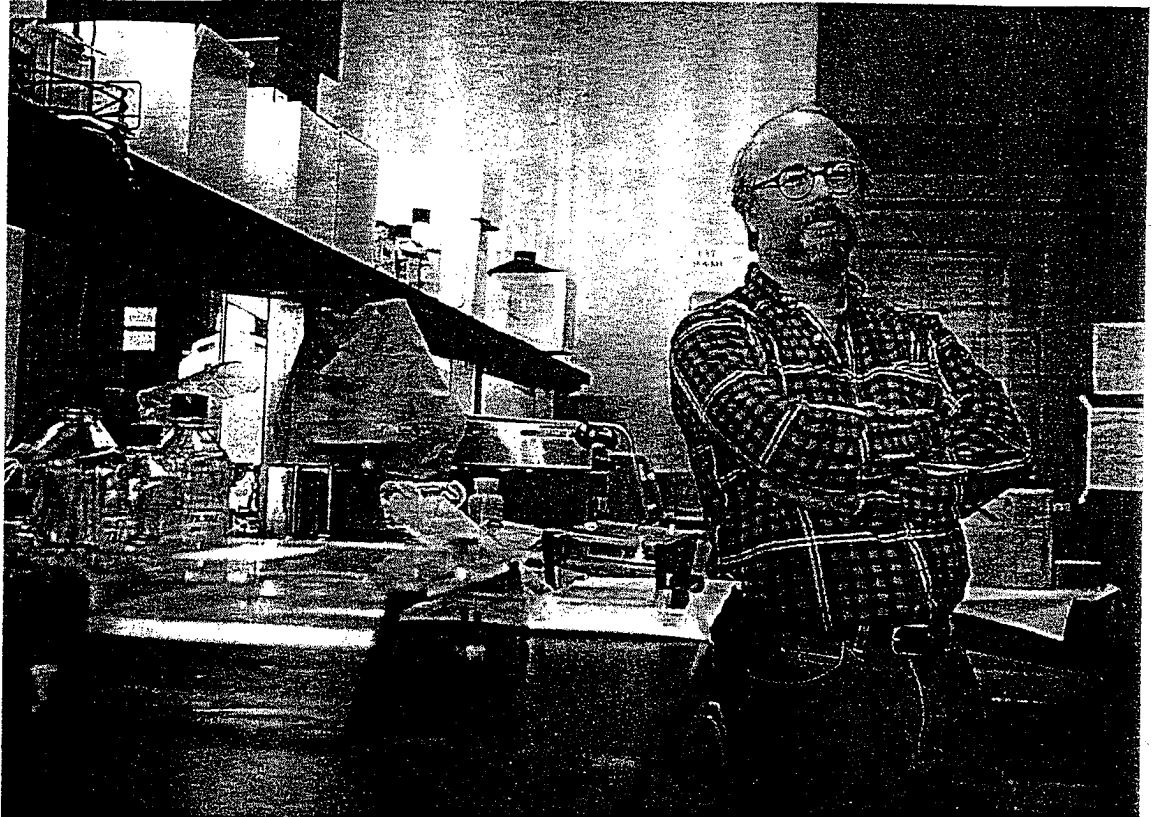
But they dispute the idea that natural selection, the force Darwin suggested drove evolution, is enough to explain the complexity of the earth's plants and animals. That complexity, they say, must be the work of an intelligent designer.

This designer may be much like the biblical God, proponents say, but they are open to other explanations, such as the proposition that life was seeded by a meteorite from elsewhere in the cosmos, possibly involving extraterrestrial intelligence, or the new age philosophy that the universe is suffused with a mysterious but inanimate life force.

In recent months, the proponents of intelligent design have advanced their case on several fronts.

In Kansas, after the backlash against the traditional biblical creationism, proponents of the design theory have become the dominant anti-evolution force, though they lost an effort to have theories like intelligent design considered on an equal basis with evolution in school curriculums.

In Michigan, nine legislators in the House of Representatives have introduced legislation to amend state education standards to put intelligent design on an equal basis with evolution.



Salvatore C. DiMarco Jr. for The New York Times

An originator of a theory on life that challenges Darwin's theory of evolution, Dr. Michael J. Behe argues that various biochemical structures in cells could not have been built in a stepwise Darwinian fashion.

In Pennsylvania, where biblical creationists and design theorists have operated in concert, state officials are close to adopting educational standards that would allow the teaching of theories on the origin and development of life other than evolution.

Backers of intelligent design organized university-sanctioned conferences at Yale and Baylor last year, and the movement has spawned at least one university student organization — called Intelligent Design and Evolution Awareness, or the IDEA club — at the University of California in San Diego.

The Discovery Institute, a research institute in Seattle that promotes conservative causes, organized a briefing on intelligent design last year on Capitol Hill for prominent members of Congress.

"They are skilled in analyzing evidence and ideas," said Representative Tom Petri, a Wisconsin Republican and one of several members of Congress who was a host at the session in a Congressional hearing room. "They are making a determined effort to attempt to present the intelligent design theory, and ask that it be judged by normal scientific criteria."

Polls show that the percentage of Americans who say they believe in creationism is about 45 percent. George W. Bush took the position in the presidential campaign that children should be exposed to both creationism and evolution in school.

Supporters of Darwin see intelligent design as more insidious than creationism, especially given that many of its advocates have mainstream scientific credentials, which creationists often lack.

"The most striking thing about the intelligent design folks is their potential to really make anti-evolutionism intellectually respectable," said Dr. Eugenie Scott, executive director of the National Center for Science Education in Oakland, Calif., which promotes the teaching of evolution.

Dr. Adrian Melott, a professor of physics and astronomy at the University of Kansas in Lawrence and a member of Kansas Citizens for Science, a group that helped win the restoration of evolution to the state education standards, said the design theory was finding adherents among doctors, engineers and people with degrees in the humanities.

Intelligent design is "the language that the creationists among the student body tend to use now," Dr. Melott said.

One of the first arguments for the

tion," Dr. Orr said.

Exactly how a designer might have assembled cell structures, say, is a question seldom addressed by design theorists. But they point out that Darwinists cannot necessarily offer detailed, step-by-step sequences of events for them either.

Dr. Behe, Dr. Dembski and Phillip E. Johnson, a professor emeritus of the law school at the University of California at Berkeley, are regarded as the intellectual fathers of the design theory movement. Mr. Johnson's book "Darwin on Trial" (InterVarsity Press, 1991) has become its manifesto. The book focuses on what Mr. Johnson says are the difficulties

Darwinian theory has in explaining the fossil record.

Until last fall, Dr. Dembski was the director of a center at Baylor that was dedicated to the study of intelligent design theory. After complaints from other Baylor faculty members, the center's focus and leadership were changed, and it now includes design theory as well as other philosophical, theological and scientific topics.

Dr. Dembski and Dr. Behe are fellows of the Discovery Institute, the Seattle research institute that promotes intelligent design in its Center for the Renewal of Science and Culture.

The center's \$1.1 million annual budget is supplied largely by Christian foundations that broadly endorse the implications of the intelligent design theory, said Bruce Chapman, Discovery's president. Mr. Johnson is an adviser to the institute, he said.

The center, which reaches people through books, articles, lectures and local activism, "is going to be of interest to academics," Mr. Chapman said. "But it's also going to be of interest to people in a more grass-roots situation because they're teaching science or because they're on a school board somewhere."

Self-Organization and Irreducibly Complex Systems: A Reply to Shanks and Joplin*

Michael J. Behe†

Department of Biological Sciences, Lehigh University

Some biochemical systems require multiple, well-matched parts in order to function, and the removal of any of the parts eliminates the function. I have previously labeled such systems “irreducibly complex,” and argued that they are stumbling blocks for Darwinian theory. Instead I proposed that they are best explained as the result of deliberate intelligent design. In a recent article Shanks and Joplin analyze and find wanting the use of irreducible complexity as a marker for intelligent design. Their primary counterexample is the Belousov-Zhabotinsky reaction, a self-organizing system in which competing reaction pathways result in a chemical oscillator. In place of irreducible complexity they offer the idea of “redundant complexity,” meaning that biochemical pathways overlap so that a loss of one or even several components can be accommodated without complete loss of function. Here I note that complexity is a quantitative property, so that conclusions we draw will be affected by how well-matched the components of a system are. I also show that not all biochemical systems are redundant. The origin of non-redundant systems requires a different explanation than redundant ones.

1. Introduction. In the past half-century biology has made astonishing progress in understanding the molecular and cellular basis of life. In light of this progress it is fair to ask whether Darwin’s mechanism of natural selection acting on random variation appears to be a good explanation for the origin of all, or just some, of the molecular systems science has discovered. In *Darwin’s Black Box: The Biochemical Challenge to Evolution* (Behe 1996) I argued that some biochemical systems, such as the blood clotting cascade or bacterial flagellum, are resistant to Darwinian expla-

*Received August 1999; revised November 1999.

†Send requests for reprints to the author, Department of Biological Sciences, Lehigh University, 111 Research Drive, Bethlehem, PA 18015.

Philosophy of Science, 67 (March 2000) pp. 155–162. 0031-8248/2000/6701-0008\$2.00
Copyright 2000 by the Philosophy of Science Association. All rights reserved.

nation because they are irreducibly complex. I defined irreducible complexity as

a single system which is composed of several well-matched, interacting parts that contribute to the basic function, and where the removal of any one of the parts causes the system to effectively cease functioning. (1996, 39)

The difficulty for Darwinian theory is that

An irreducibly complex system cannot be produced directly (that is, by continuously improving the initial function, which continues to work by the same mechanism) by slight, successive modifications of a precursor system, because any precursor to an irreducibly complex system that is missing a part is by definition nonfunctional. (1996, 39)

To illustrate the concept with a familiar example for a general readership, I pointed to a simple mechanical mousetrap, composed of several parts such as the base, hammer, spring, and so on, and noted that the absence of any of the parts destroys the mouse-catching ability of the trap. Darwin's vision of natural selection gradually improving function in "numerous, successive, slight modifications" (Darwin 1859) appears not to fit well with such systems. I went on to argue that, since intelligent agents are the only entities known to be able to construct irreducibly complex systems, the biochemical systems are better explained as the result of deliberate intelligent design.

But are gradual Darwinian natural selection and intelligent design the only potential explanations? Shanks and Joplin (1999) direct our attention to complexity theory, which concerns the ability of systems to self-organize abruptly, sometimes in surprising ways. They suggest that irreducibly complex biochemical systems might in principle be explained by self-organization, eliminating the need to invoke intelligence. They then go on to argue that biochemical systems are "redundantly complex"—that is, contain components that can be removed without entirely eliminating function.

After briefly describing the Belousov-Zhabotinsky reaction—Shanks and Joplin's main counterexample—I will first argue that the reaction does not meet the definition of irreducibly complex, because the interacting components are not "well-matched." I will then agree that redundant complexity exists, but show that not all of biochemistry is redundant.

2. A Closer Look At Chemical Self-Organization. The dissipation of energy in nature can organize matter and produce reaction pathways. A simple example is the clumping of matter into stars under the influence of gravity. More complex examples are tornados and the stellar nuclear pathways

that lead to the production of the heavy elements. These examples, however, have no direct relevance to the origin of biochemical systems. Shanks and Joplin (1999) offer what they think is a more pertinent example—the Belousov-Zhabotinsky reaction, a self-organizing chemical system discovered in the 1950s by B. P. Belousov in an attempt to model the Krebs cycle. The term “BZ reaction” is applied to a group of chemical reactions in which an organic substrate is oxidized by bromate ions in the presence of a transition metal ion and acid. Instead of proceeding monotonically to equilibrium, the reaction oscillates between two pathways because of a competition between bromide ion and bromous acid for reaction with bromate ion. Bromate oxidizes the metal ions, which in turn are re-reduced by reaction with organic substrate. When the reaction is well-stirred, the visible result is a solution that switches from one color to another at constant time intervals until the reaction materials are consumed. When the same reaction is set up as a thin, unstirred layer, waves of color change propagate through the layer. For details of the reaction pathways, see Gray and Scott 1994 and references therein.

Shanks and Joplin write that the BZ reaction “satisfies Behe’s criteria for irreducible chemical complexity” because if any of the chemical components is removed “the characteristic behavior of the system is disrupted.” Thus “Irreducible complexity in a self-organizing system” can be generated “without the aid of a designing deus ex machina” (1999, 272–273).

I disagree that the BZ reaction “satisfies Behe’s criteria” for an irreducibly complex system. Although it does have interacting parts that are required for the reaction, the system lacks a crucial feature—the components are not “well-matched.” The appearance of the modifier “well-matched” in the definition I constructed (above) reflects the fact that complexity is a quantitative property. A system can be more or less complex, so the likelihood of coming up with any particular interactive system by chance can be more or less probable. As an illustration, contrast the greater complexity of a mechanical mousetrap (mentioned above) with the much lesser complexity of a lever and fulcrum. Together a lever and fulcrum form an interactive system which can be used to move weights. Nonetheless, the parts of the system can have a wide variety of shapes and sizes and still function. Because the system is not well-matched, it could easily be formed by chance.

Systems requiring several parts to function that need not be well-matched, we can call “simple interactive” systems (designated ‘SI’). Ones that require well-matched components are irreducibly complex (‘IC’). The line dividing SI and IC systems is not sharp, because assignment to one or the other category is based on probabilistic factors which often are hard to calculate and generally have to be intuitively estimated based on always-incomplete background knowledge. Moreover, no law of physics auto-

matically rules out the chance origin of even the most intricate IC system. As complexity increases, however, the odds become so abysmally low that we reject chance as an explanation (Dembski 1998).

Just as I think that a gradual origin by natural selection is a good explanation for some things, I agree that a discontinuous origin by self-organization explains some things too. Nonetheless, I do not think either explains irreducible complexity. I argue that Shanks and Joplin's counterexample—the BZ reaction—is not IC; it is SI, because the components are not well-matched. To justify my position, let me first illustrate a well-matched system using the blood clotting cascade (Stubbs and Bode 1994). The active form of one protein of the cascade is called thrombin, which cleaves the soluble protein fibrinogen to produce fibrin, the insoluble meshwork of a blood clot. The chemistry catalyzed by thrombin is simply the hydrolysis of a certain fibrinogen peptide bond. However, all proteins are made of amino acid residues joined by peptide bonds. A typical protein contains several hundred peptide bonds. There is nothing remarkable about the bond in fibrinogen that is cleaved by thrombin. Yet thrombin selects that particular bond for cleavage out of literally hundreds of thousands of peptide bonds in its environment and ignores almost all others. It can do this because the shape of thrombin is well-matched to the shape of fibrinogen around the bond it cleaves. It “recognizes” not only the bond it cuts, but also a number of other features of its target. The other proteins of the clotting cascade (Stuart factor, proaccelerin, tissue factor, and so on) have similar powers of discrimination. So do virtually all of the components of the molecular machines I discussed in *Darwin's Black Box*.

Let us contrast this biochemical specificity with a comparable chemical reaction lacking such specificity. The peptide bonds of proteins can also be cleaved by simple chemicals. A typical procedure calls for incubating the protein in 6*N* hydrochloric acid at 110°C for twenty four hours. If fibrinogen were incubated under those conditions, the peptide bond that thrombin cleaves would be broken, but so would every other peptide bond in the protein. It would be completely reduced to amino acids. If thrombin were in the mix, it too would be completely destroyed. If the other proteins of the clotting cascade were there, no clotting would take place, even though the peptide bonds that are cleaved in the cascade would be cleaved, because all other peptide bonds would be hydrolyzed too. There is virtually no specificity to the chemical hydrolysis beyond the type of bond that is cleaved.

Similarly, the reactants of the BZ reaction are small organic or inorganic chemicals that show little specificity for each other. One ingredient, sodium bromate, is a general purpose oxidizing reagent and is capable of degrading a very large spectrum of chemicals besides the ones used in BZ reactions (thus its transport aboard airlines is forbidden). Another re-

quirement of the reaction is simply for a transition metal that can change its oxidation state, and a number of such metals are known, including iron, cerium, and manganese ions (Field 1972). A third requirement is for an organic molecule that can be oxidized. Many candidates could fulfill this role (ones that have been used include malonic, citric, maleic, and malic acids), and organic molecules can be oxidized by many reagents other than bromate. The last ingredient is simply a high concentration of sulfuric acid. As Field (1972, 308) noted, setting up BZ reactions "is an exceedingly easy task as they will occur over a wide range of concentrations and conditions."

The BZ class of self-organizing reactions—chemical oscillations—is surprising and interesting. Nonetheless, its complexity can be likened to other self-organizing systems found outside of biology, such as, say, tornados, which, although they command our attention, do not approach the specificity of well-matched, irreducibly complex biochemical systems.

3. Biochemical Self-Organization: Behavior vs. Origin. The dynamical behavior of the BZ reaction has been modeled by a set of two ordinary differential equations (Tyson 1994, 577). Because some biological systems can be modeled by similar mathematics, Shanks and Joplin (1999) conclude that self-organization can explain the behavior of the biological systems. There are several reasons to question the relevance of their point. First, they also note that "the substrates and products in these systems are very different from those in the BZ reaction" (1999, 273). In other words, we have traveled far from cerium, sodium bromate, and the other constituents of the chemical system. Second, and more importantly, the behavior of a system must be distinguished from its origin. As an illustration, consider highway traffic flow. A number of mathematical models have been used to describe traffic flow, some drawing on theories of self-organization (Schreckenberg and Wolf 1998). The mathematics, however, have not called the automobiles into being. The mathematics simply try to describe the typical behavior of traffic when a certain density is reached under conditions of restricted movement on a highway.

Examples of biological processes that show BZ dynamical behavior include glycolysis and aggregation of dispersed cells of the slime mold *Dictyostelium discoideum* into a slug. But consider the sophisticated components of the aggregation-signaling system of *D. discoideum*, which include: a cyclic AMP membrane receptor protein that can exist in an active and inactive form; an adenylate cyclase that binds to the active form of the receptor and itself becomes activated; a protein to export cyclic AMP into the extracellular medium; and more (Goldbeter 1996, Part I). All of that complicated machinery is ignored in BZ models—treated as a black box. Oscillations in the cellular concentration of glycolytic intermediates

are due in large part to the multi-talented phosphofructokinase (PFK), a tetrameric enzyme that can exist in two conformational states (an active form and a less-active one) and which has binding sites for a dozen different activators and inhibitors (Goldbeter 1996, Part III). Mathematical models of BZ behavior do not explain the origin of the impressive abilities of PFK any more than models of traffic flow explain the origin of brakes or gas pedals. Thus, even if a biological system displays self-organizing behavior, the question of its origin remains.

4. Not All Biochemical Systems Are Redundant. In contrast to claims about irreducible complexity, Shanks and Joplin write that “Real biochemical systems, we argue, manifest *redundant complexity*—a characteristic result of evolutionary processes” (1999, 268). By this they mean that biochemical pathways overlap and are interconnected, so that removal of one or even several components does not completely destroy the function. In support of their position they cite a diverse array of biochemical examples: the synthesis of an alternate pine tree lignin with increased content of dihydroconiferyl alcohol; viable mice in which the gene for the tumor suppressor *p53* was knocked out; and more. Their initial illustration is the metabolic pathways for the synthesis of glucose-6-phosphate. They point out that the molecule can be made by “several different isoforms or variants of hexokinase, and all are present, as a result of gene duplication, in varying proportions in different tissues.” What’s more, “Knock out one enzyme isoform and the other isoforms in the tissue can take over its function” (277).

True enough. The observation that some biochemical systems are redundant, however, does not entail that all are. And, in fact, some are not redundant. Consider the following examples of nonredundant metabolic pathways. Primates, including humans, cannot synthesize ascorbic acid (vitamin C) because they lack a functional gene for L-gulonogamma-lactone oxidase, although a pseudogene is present (Nishikimi and Yagi 1991). Vitamin C is made by no other pathway. Hexosaminidase A is required to catabolize ganglioside G_{M2} ; its loss results in Tay-Sachs disease (Kolter and Sandhoff 1998). These enzymes are parts of “real biochemical systems,” but they do not “manifest redundant complexity.” (For many, many additional examples, see Scriver 1995 or other texts on inborn errors of metabolism.) Therefore, arguments developed about the origin of redundant systems do not necessarily apply to all biochemical systems.

Shanks and Joplin’s argument for redundant complexity has the same strengths and weaknesses when the subject moves from metabolic pathways to other biochemical systems. That is, they are right to notice that some systems or components are redundant, but wrong to extrapolate the conclusion to all systems. For example, they point to mice in which the